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NEWS 13 MAY 14 DGENE, PCTGEN and USGENE enhanced with increased
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                introduction of free HIT display format
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                status data
NEWS 15 MAY 28 CAS databases on STN enhanced with NANO super role in
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                enhanced on STN
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```
chain nodes :
25 26 27 30
ring nodes :
1 2 3 4 5 6 7 8 9 10 11 12 13 14 15 16 17 18 19 20 21 22 23
24
chain bonds :
6-25 10-27 14-26 17-20 25-27 26-27
ring bonds :
1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12 13-14 13-18
14-15 15-16 16-17 17-18 19-20 19-24 20-21 21-22 22-23 23-24
exact/norm bonds :
6-25 10-27 13-14 13-18 14-15 14-26 15-16 16-17 17-18 17-20 25-27 26-27
normalized bonds :
1-2 1-6 2-3 3-4 4-5 5-6 7-8 7-12 8-9 9-10 10-11 11-12 19-20 19-24
20-21 21-22 22-23 23-24
isolated ring systems :
containing 1 : 7 : 13 : 19 :
```

### G1:C,N

G2:CF2,CF3,CC12,CC13,CBr2,CBr3,X

# G3:C,N

Match level :

1:Atom 2:Atom 3:Atom 4:Atom 5:Atom 6:Atom 7:Atom 8:Atom 9:Atom 10:Atom 11:Atom 12:Atom 13:Atom 14:Atom 15:Atom 16:Atom 17:Atom 18:Atom 19:Atom 20:Atom 21:Atom 22:Atom 23:Atom 24:Atom 25:CLASS 26:CLASS 27:CLASS 30:CLASS 31:Atom

L1 STRUCTURE UPLOADED

=> d 11

L1 HAS NO ANSWERS

L1

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=> s 11 full

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FULL SCREEN SEARCH COMPLETED -20268 TO ITERATE

100.0% PROCESSED 20268 ITERATIONS SEARCH TIME: 00.00.02

211 ANSWERS

186.10

211 SEA SSS FUL L1

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COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION 185.88

FULL ESTIMATED COST

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L3 ANSWER 1 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2009:412461 CAPLUS

DOCUMENT NUMBER: 151:496

TITLE: QSAR study of the 5-HT1A receptor affinities of

arylpiperazines using a genetic algorithm-artificial neural network model

AUTHOR(S): Habibi-Yangjeh, Aziz

CORPORATE SOURCE: Department of Chemistry, Faculty of Science,

University of Mohaghegh Ardabili, Ardabil, Iran SOURCE: Monatshefte fuer Chemie (2009), 140(5), 523-530

CODEN: MOCMB7; ISSN: 0026-9247

PUBLISHER: SpringerWienNewYork

DOCUMENT TYPE: Journal LANGUAGE: English

AB Genetic algorithm-multiparameter linear regression (GA-MLR) and genetic algorithm-artificial neural network (GA-ANN) models have been used for prediction of the 5-HTIA receptor affinities (pK i) of 66 arylpiperaxines. A large number of theor. descriptors were calculated for each compound The cenetic

algorithm (GA) was used for selection of the variables that resulted in the best fit to the MLR and ANN models. The models were generated using seven descriptors as variables. For evaluation of the predictive power of the models, pK i values of 13 compds. in the prediction set were calculated Mean percentage deviation (MPD) for the GA-MLR and GA-ANN models were 0.344 and 0.065, resp. Comparison of the results obtained by use of the models reveals the GA-ANN model is superior to the GA-MLR model. Graphical abstract

IT 328248-21-1 328248-24-4 328248-30-2 328248-36-8 753439-74-6 767277-20-3

777843-82-0

RL: PAC (Pharmacological activity); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(QSAR study of 5-HT1A receptor affinities of arylpiperazines using a genetic algorithm-artificial neural network model)

RN 328248-21-1 CAPLUS

CN Piperazine, 1-(4-methoxyphenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

RN 328248-24-4 CAPLUS

CN Piperazine, 1-(2-chlorophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

$$\begin{array}{c} \text{C1} & \text{N---} \text{CH}_2\text{---} \text{CH---} \text{O} \\ \end{array}$$

RN 328248-30-2 CAPLUS

CN Piperazine, 1-(4-nitrophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

RN 328248-36-8 CAPLUS

CN Piperazine, 1-(2-methoxyphenyl)-4-[3-(4-methoxyphenyl)-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

RN 753439-74-6 CAPLUS

CN Piperazine, 1-(4-fluorophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

$$\begin{array}{c} \text{Ph} \\ \text{N---} \text{CH}_2\text{---} \text{CH}_2\text{---} \text{CH---} \text{O} \end{array}$$

RN 767277-20-3 CAPLUS

CN Piperazine, 1-[3-[1,1'-biphenyl]-4-yl-3-[4-(trifluoromethyl)phenoxy]propyl]-4-(2-methoxyphenyl)- (CA INDEX NAME)

RN 777843-82-0 CAPLUS

CN Piperazine, 1-(2-methoxyphenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

$$\begin{array}{c|c} \text{OMe} & \text{Ph} & \text{CF}; \\ \hline \text{OMe} & \text{N} - \text{CH}_2 - \text{CH}_2 - \text{CH} - \text{O} \\ \end{array}$$

REFERENCE COUNT:

46 THERE ARE 46 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 2 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2008:986436 CAPLUS

DOCUMENT NUMBER: 150:321888

TITLE: The structure-based 3D-QSAR study of MCH1 receptor

antagonists

AUTHOR(S): Lee, New Gil; Yoo, Seung-Eun; Kang, Nam Sook
CORPORATE SOURCE: Korea Research Institute of Chemical Technology,

Daejeon, S. Korea
SOURCE: Molecular Simulation (2008), 34(7), 699-705

CODEN: MOSIEA: ISSN: 0892-7022

PUBLISHER: Taylor & Francis Ltd.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB Melanin-concentrating hormone 1 receptor (MCH1-R) mediates the orexigenic effects

of melanin-concentrating hormone and its antagonist, are considered as a potential targets for the treatment of obesity. To design more potent and selective MCHI-R antagonists, at first, the authors built up the homol. structure of MCHI-R. Then, the authors carried out the receptor based 3 dimensional Quant. Structure Activity Relationship (3D-QSAR) using comparative mol. field anal. and Comparative Mol. Similarity Indexes Anal. (CCMSIA) for a series of scaffold of MCHI-R antagonists and the docking study for MCHI-R. These models are proved as statistically valid models with a good correlative and predictive power. Based on these models, the authors are going to develop more potent and selective MCHI-R antagonists. 1132777-99-1

1132///-99-1

RL: PEP (Physical, engineering or chemical process); PRP (Properties); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (structure-based 3D-QSAR study of MCH1 receptor antagonists as anti-obesity drugs)

RN 1132777-99-1 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-bromophenoxy)-3-phenylpropy1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

REFERENCE COUNT:

23 THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 3 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2008:803320 CAPLUS

DOCUMENT NUMBER: 149:215113

TITLE: Two-dimensional QSAR studies on arylpiperazines as

high-affinity 5-HT1A receptor ligands AUTHOR(S): Weber, Karen C.; Honorio, Kathia M.; Andricopulo,

Adriano D.; Da Silva, Alberico B. F.

CORPORATE SOURCE: Instituto de Ouimica de Sao Carlos, Universidade de

Sao Paulo, Sao Carlos, 13560-970, Brazil SOURCE: Medicinal Chemistry (2008), 4(4), 328-335

CODEN: MCEHAJ; ISSN: 1573-4064

PUBLISHER: Bentham Science Publishers Ltd.

DOCUMENT TYPE: Journal English

LANGUAGE:

- 5-HT1A receptor plays an important role in the delayed onset of antidepressant action of a class of selective serotonin reuptake inhibitors. Moreover, 5-HT1A receptor levels have been shown to be altered in patients suffering from major depression. In this work, hologram quant. structure-activity relationship (HOSAR) studies were performed on a series of arylpiperazine compds, presenting affinity to the 5-HT1A receptor. The models were constructed with a training set of 70 compds. The most significant HQSAR model (q2 = 0.81, r2 = 0.96) was generated using atoms, bonds, connections, chirality, and donor and acceptor as fragment distinction, with fragment size of 6-9. Predictions for an external test set containing 20 compds. are in good agreement with exptl. results showing the robustness of the model. Addnl., useful information can be obtained from the 2D contribution maps.
- 328248-15-3 328248-21-1 328248-24-4 328248-30-2 328248-36-8 753439-74-6

767277-20-3 777843-82-0

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(two-dimensional QSAR studies on arylpiperazines as high-affinity 5-HT1A receptor ligands)

RN 328248-15-3 CAPLUS

CN Piperazine, 1-(4-chlorophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy[propyl]- (CA INDEX NAME)

RN 328248-21-1 CAPLUS

CN Piperazine, 1-(4-methoxyphenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy[propyl]- (CA INDEX NAME)

RN 328248-24-4 CAPLUS

CN Piperazine, 1-(2-chloropheny1)-4-[3-pheny1-3-[4-(trifluoromethy1)phenoxy]propy1]- (CA INDEX NAME)

RN 328248-30-2 CAPLUS

CN Piperazine, 1-(4-nitrophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

$$\mathsf{O_{2}N} \qquad \mathsf{N-CH_{2}-CH_{2}-CH-O} \qquad \mathsf{CF}$$

RN 328248-36-8 CAPLUS

CN Piperazine, 1-(2-methoxyphenyl)-4-[3-(4-methoxyphenyl)-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

RN 753439-74-6 CAPLUS

CN Piperazine, 1-(4-fluorophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

RN 767277-20-3 CAPLUS

CN Piperazine, 1-[3-[1,1'-bipheny1]-4-y1-3-[4-(trifluoromethy1)phenoxy]propy1]-4-(2-methoxypheny1)- (CA INDEX NAME)

- RN 777843-82-0 CAPLUS
- CN Piperazine, 1-(2-methoxyphenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

$$\begin{array}{c|c} \text{OMe} & \text{Ph} & \text{CF} \\ \text{N} & \text{CH}_2\text{-CH}_2\text{-CH-O} \end{array}$$

REFERENCE COUNT:

31 THERE ARE 31 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 4 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2008:767635 CAPLUS

DOCUMENT NUMBER: 149:324283

TITLE: Quantitative structure-affinity relationship of 5-HT1A

receptor ligands by the classification tree method AUTHOR(S): Kuz'min, V. E.; Polischuk, P. G.; Artemenko, A. G.;

Makan, S. Yu.; Andronati, S. A.

CORPORATE SOURCE: A.V. Bogatsky Physical-Chemical Institute, National
Academy of Sciences of Ukraine, Odessa, Ukraine

SOURCE: SAR and QSAR in Environmental Research (2008),

19(3-4), 213-244

CODEN: SQERED; ISSN: 1062-936X
PUBLISHER: Taylor & Francis Ltd.

PUBLISHER: Taylor & Francis Ltd.
DOCUMENT TYPE: Journal

DOCUMENT TYPE: Journal LANGUAGE: English

AB The influence of mol. structure of 346 ligands on their affinity for 5-HTIA receptors was investigated. It was shown that the effectiveness of the proposed novel approach for interpretation of decision tree models compared favorably with the PLS method. In the context of the proposed approach, mol. fragments and their values of the relative influence on the affinity for 5-HTIA receptors were defined.

328248-15-3 328248-21-1 328248-24-4 328248-30-2 328248-36-8 753439-74-6

767277-20-3 777843-82-0

RL: ANT (Analyte); BSU (Biological study, unclassified); PRP (Properties); ANST (Analytical study); BIOL (Biological study)

(quant. structure-affinity relationship of 5-HT1A receptor ligands by the classification tree method)

RN 328248-15-3 CAPLUS

CN Piperazine, 1-(4-chlorophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxylpropyl]- (CA INDEX NAME)

RN 328248-21-1 CAPLUS

CN Piperazine, 1-(4-methoxyphenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

$$\stackrel{\text{MeO}}{\underset{N_{\perp}}{\bigvee}} \stackrel{\text{N}}{\underset{N_{\perp}}{\bigvee}} \operatorname{CH}_2 - \operatorname{CH}_2 - \operatorname{CH}_0$$

RN 328248-24-4 CAPLUS

CN Piperazine, 1-(2-chlorophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy|propyl]- (CA INDEX NAME)

RN 328248-30-2 CAPLUS

CN Piperazine, 1-(4-nitrophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

RN 328248-36-8 CAPLUS

CN Piperazine, 1-(2-methoxyphenyl)-4-[3-(4-methoxyphenyl)-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

RN 753439-74-6 CAPLUS

CN Piperazine, 1-(4-fluorophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

$$\begin{array}{c} Ph \\ N-CH_2-CH_2-CH-O \end{array}$$

RN 767277-20-3 CAPLUS

CN Piperazine, 1-[3-[1,1'-bipheny1]-4-y1-3-[4-(trifluoromethyl)phenoxy]propy1]-4-(2-methoxypheny1)- (CA INDEX NAME)

RN 777843-82-0 CAPLUS

CN Piperazine, 1-(2-methoxyphenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

$$\begin{array}{c|c} \text{OMe} & \text{Ph} & \text{CF} \\ \hline \text{N} & \text{CH}_2\text{-CH}_2\text{-CH}-\text{O} \end{array}$$

REFERENCE COUNT:

25 THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 5 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2008:232006 CAPLUS

DOCUMENT NUMBER: 148:440268

TITLE: A chemometric study of the 5-HT1A receptor affinities

presented by arylpiperazine compounds
AUTHOR(S): Weber, Karen C.; da Silva, Alberico B. F.

CORPORATE SOURCE: Instituto de Quimica de Sao Carlos, Universidade de

Sao Paulo, Sao Carlos, 13566-590, Brazil
SOURCE: European Journal of Medicinal Chemistry (2008), 43(2),

364-372

CODEN: EJMCA5; ISSN: 0223-5234

PUBLISHER: Elsevier Masson SAS DOCUMENT TYPE: Journal

DOCUMENT TYPE: Journal LANGUAGE: English

AB Arylpiperazine compds. are promising 5-HT1A receptor ligands that can contribute for accelerating the onset of therapeutic effect of selective serotonin reuptake inhibitors. In the present work, the chemometric methods EGA, PCA, KNN, SINCA and PLS were employed in order to obtain SAR and QSAR models relating the structures of arylpiperazine compds. to their 5-HT1A receptor affinities. A training set of 52 compds. was used to construct the models and the best ones were obtained with nine topol. descriptors. The classification and regression models were externally validated by means of predictions for a test set of 14 compds. and have presented good quality, as verified by the correctness of classifications, in the case of pattern recognition studies, and by the high correlation coeffs. (q2 = 0.76, r2 = 0.83) and small prediction errors for the PLS regression. Since the results are in good agreement with previous SAR studies, we can suggest that these findings can help in the search for 5-HT1A receptor liquands that are able to improve antidepressant treatment.

IT 328248-21-1 328248-24-4 328248-30-2 328248-36-8 753439-74-6 767277-20-3

777843-82-0

RL: DMA (Drug mechanism of action); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(chemometric study of 5-HT1A receptor affinities presented by arylpiperazine compds. as possible antidepressants)

RN 328248-21-1 CAPLUS

CN

Piperazine, 1-(4-methoxyphenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

RN 328248-24-4 CAPLUS

CN Piperazine, 1-(2-chlorophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxylpropyl]- (CA INDEX NAME)

$$\begin{array}{c} \text{C1} & \text{Ph} \\ \text{N---} \text{CH}_2\text{---} \text{CH}_2\text{---} \text{CH---} \text{O} \end{array}$$

RN 328248-30-2 CAPLUS

CN Piperazine, 1-(4-nitropheny1)-4-[3-pheny1-3-[4-(trifluoromethy1)phenoxy]propy1]- (CA INDEX NAME)

RN 328248-36-8 CAPLUS

CN Piperazine, 1-(2-methoxyphenyl)-4-[3-(4-methoxyphenyl)-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

RN 753439-74-6 CAPLUS

CN Piperazine, 1-(4-fluorophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

$$\begin{array}{c} \text{F} \\ \text{N} \\ \text{CH}_2 \\ \text{CH}_2 \\ \text{CH}_2 \\ \text{CH} \\ \text{O} \end{array}$$

RN 767277-20-3 CAPLUS

CN Piperazine, 1=[3-[1,1'-bipheny1]-4-y1-3-[4-(trifluoromethyl)phenoxy]propy1]-4-(2-methoxypheny1)- (CA INDEX NAME)

RN 777843-82-0 CAPLUS

CN Piperazine, 1-(2-methoxyphenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

$$\begin{array}{c|c} \text{OMe} & \text{Ph} & \text{CF} \\ \hline \text{N} & \text{CH}_2\text{-CH}_2\text{-CH}-\text{O} \end{array}$$

REFERENCE COUNT:

35 THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 6 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2006:847178 CAPLUS

DOCUMENT NUMBER: 145:410017

TITLE: Synthesis of benzenepropanamine analogues as non-detergent spermicides with antitrichomonas and

anticandida activities

AUTHOR(S): Kumar, S. T. V. S. Kiran; Sharma, Vishnu Lal; Kumar,
Manish; Shukla, Praveen Kumar; Tiwari, Pratibha; Jain,

Rajeev Kumar; Maikhuri, Jagdamba Prasad; Singh, Divya; Gupta, Gopal; Singh, Man Mohan

CORPORATE SOURCE: Division of Medicinal and Process Chemistry, Central Drug Research Institute, Lucknow, 226001, India

SOURCE: Drug Research Institute, Lucknow, 226001, India Source: Bioorganic & Medicinal Chemistry (2006), 14(19),

6593-6600

CODEN: BMECEP; ISSN: 0968-0896

PUBLISHER: Elsevier Ltd.
DOCUMENT TYPE: Journal
LANGUAGE: English

LANGUAGE: English
OTHER SOURCE(S): CASREACT 145:410017

GI

Ι

AB Fifteen analogs of benzenepropanamine were synthesized and evaluated for their spermicidal as well as microbicidal activities against Trichomonas vaginalis and Candida spp. Several compds. showed appreciable dual activities. Compound I exhibited good spermicidal (MEC = 0.1%) along with substantial anticandidal (MIC = 0.05%) activities, while compds. 3 and 6 showed significant microbicidal activities with moderate spermicidal effect. The SAR of these structures is being discussed here in this communication. It is concluded that suitable structural modifications in this class of compds. at 3-amino position may lead to a potent spermicide with associated microbicidal activity.

IT 911811-08-0P 911811-09-1P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(benzenepropanamine analogs as non-detergent spermicides with antitrichomonas and anticandida activities)

- RN 911811-08-0 CAPLUS
- CN Piperazine, 1-(3-chlorophenyl)-4-[3-phenyl-3-[4-

(trifluoromethyl)phenoxy[propyl]-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

- RN 911811-09-1 CAPLUS
- CN Piperazine, 1-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]-4-[3-(trifluoromethyl)phenyl]-, hydrochloride (1:1) (CA INDEX NAME)

$$\begin{array}{c} \text{Ph} \\ \text{F}_{3}\text{C} \\ \end{array}$$

● HCl

REFERENCE COUNT:

34 THERE ARE 34 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 7 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN 1.3

ACCESSION NUMBER: 2005:1143268 CAPLUS

DOCUMENT NUMBER: 144:63874

TITLE: Design and synthesis of long-chain arylpiperazines with mixed affinity for serotonin transporter (SERT)

and 5-HT1A receptor

Perrone, Roberto; Berardi, Francesco; Colabufo, Nicola AUTHOR(S):

A.; Lacivita, Enza; Larizza, Carmela; Leopoldo,

Marcello: Tortorella, Vincenzo

CORPORATE SOURCE: Dipartimento Farmaco-Chimico, Universita degli Studi

di Bari, Bari, 70125, Italy

SOURCE: Journal of Pharmacy and Pharmacology (2005), 57(10), 1319-1327

CODEN: JPPMAB; ISSN: 0022-3573 PUBLISHER . Pharmaceutical Press

DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 144:63874

A new generation of antidepressant agents could be represented by compds. with mixed activity as serotonin transporter (SERT) inhibitors and 5-HT1A receptor antagonists. We report here on the synthesis and evaluation of SERT and 5-HT1A receptor affinity of long-chain arylpiperazines obtained either by modifying 6-nitroquipazine into a long-chain arylpiperazine or by inserting a modified 6-nitroquipazine moiety or other structures endowed with SERT affinity into a long-chain arylpiperazine with 5-HT1A affinity. Among the compds. studied, 2-[4-(2-methoxyphenyl)piperazin-1-yl]-N-(6-nitro-2-quinolyl)ethylamine

(21) and 1-(5-bromo-1,2,3,4-tetrahydronaphthalen-1-y1)-3-[4-(2methoxyphenyl)-piperazin-1-yl]-1-propanone (24) showed good affinity values for SERT and 5-HT1A receptors (SERT: Ki (inhibition constant) = 71.8

and 62.8 nM; 5-HT1A Ki = 14.2 and 0.82 nM, resp.). 871739-17-2P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(arylpiperazines with mixed affinity for serotonin transporter and 5-HT1A receptor)

RN 871739-17-2 CAPLUS

CN Piperazine, 1-phenyl-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxylpropyl]-, hydrochloride (1:2) (CA INDEX NAME)

#### ● 2 HC1

#### 777843-82-0

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (arylpiperazines with mixed affinity for serotonin transporter and

5-HT1A receptor)

RN

777843-82-0 CAPLUS
Piperazine, 1-(2-methoxyphenyl)-4-[3-phenyl-3-[4-CN (trifluoromethyl)phenoxy[propyl]- (CA INDEX NAME)

29

REFERENCE COUNT:

THERE ARE 29 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

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L3 ANSWER 8 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2005:1103625 CAPLUS

DOCUMENT NUMBER: 143:387060

TITLE: Preparation of piperazine or piperidine derivatives as

serotonin reuptake inhibitors

INVENTOR(S): Pinney, Kevin G.; Miranda, Maria Graciela; Dorsey,

James Michael

PATENT ASSIGNEE(S): Baylor University, USA SOURCE: PCT Int. Appl., 52 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.				KIND DAT			APPLICATION NO.						DATE					
WO	WO 2005094896				A2		20051013		WO 2005-US10356					20050328				
WO	WO 2005094896				A3 2		20070503											
	W:	AE.	AG.	AL.	AM.	AT.	AU,	AZ.	BA.	BB.	BG.	BR.	BW.	BY.	BZ.	CA.	CH.	
							DE,											
							ID,											
							LV,											
							PL,											
							TT,											ZW
	RW:	BW,	GH,	GM,	KE,	LS,	MW,	ΜZ,	NA,	SD,	SL,	SZ,	TZ,	UG,	ZM,	ZW,	AM,	
		AZ,	BY,	KG,	ΚZ,	MD,	RU,	ΤJ,	TM,	AT,	BE,	BG,	CH,	CY,	CZ,	DE,	DK,	
		EE,	ES,	FI,	FR,	GB,	GR,	HU,	IE,	IS,	IT,	LT,	LU,	MC,	NL,	PL,	PT,	
		RO.	SE.	SI.	SK.	TR.	BF.	BJ.	CF.	CG.	CI.	CM.	GA.	GN.	GO.	GW.	ML.	
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								EP 2005-730778					20050328					
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	14.						MC,											
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				MK,											_			
US 20080132514																		
PRIORITY APPLN. INFO.:							US 2004-557069P				1	P 2	0040.	326				
										WO 2	005-	US10	356	1	W 2	0050	328	
OTHER SOURCE(S): CASREACT 143:387060; MARPAT 143:387060																		

Title compds. I [X = F or CF3; Y = (CH2)n; n = 0-1; A = N or C; Ar = aryl]AB and their pharmaceutically acceptable salts, are prepared and disclosed as serotonin reuptake inhibitors. Thus, e.g., II was prepared by reduction of 1-phenyl-3-(4-phenyl-piperazin-1-yl)-propan-1-ol (preparation given) using sodium borohydride followed by coupling with 4-fluorophenol. The ability of I to inhibit [3H]5-HT uptake was evaluated using liquid scintillation spectroscopy and it was revealed that selected compds. of the invention possessed IC50 values in the range of 1.45 up to 9.56 µM. I as serotonin reuptake inhibitors should prove useful in the treatment of depression. Pharmaceutical composition comprising I are disclosed.

II

TT 158545-85-8P 691872-56-7P 691872-58-9P 691872-60-3P 691872-62-5P 691872-64-7P 691872-66-9P 866548-21-2P 866548-22-3P 866548-23-4P 866548-24-5P 866548-25-6P 866548-26-7P 866548-27-8P 866548-28-9P 866548-29-0P 866548-30-3P 866548-31-4P 866548-36-9P 866548-37-0P RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(Uses) (preparation of piperazine or piperidine derivs. as serotonin reuptake inhibitors)

RN

158545-65-8 CAPLUS
Piperazine, 1-phenyl-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]-(CA INDEX NAME)

RN 691872-56-7 CAPLUS

CN Piperazine, 1-(3-chlorophenyl)-4-[2-(4-fluorophenoxy)-2-phenylethyl]- (CA INDEX NAME)

691872-58-9 CAPLUS
Piperazine, 1-[2-(4-fluorophenoxy)-2-phenylethyl]-4-(2-methoxyphenyl)-CN (CA INDEX NAME)

$$\begin{array}{c|c} \text{OMe} & \text{Ph} \\ \hline & \text{N} - \text{CH}_2 - \text{CH} - \text{O} \end{array}$$

RN 691872-60-3 CAPLUS

CN Piperazine, 1-[2-(4-fluorophenoxy)-2-phenylethyl]-4-[3-(trifluoromethyl)phenyl]- (CA INDEX NAME)

$$^{\rm F3C} \stackrel{\rm Ph}{\longrightarrow} ^{\rm N-CH_2-CH-O}$$

691872-62-5 CAPLUS RN

Piperazine, 1-(3-chlorophenyl)-4-[2-(4-fluorophenoxy)-2-phenylethyl]-, CN hydrochloride (1:1) (CA INDEX NAME)

● HC1

RN 691872-64-7 CAPLUS
CN Piperazine, 1-[2-(4-fluorophenoxy)-2-phenylethyl]-4-(2-methoxyphenyl)-,
hydrochloride (1:1) (CA INDEX NAME)

$$\stackrel{\text{OMe}}{\underset{N}{\longrightarrow}} \text{N---} \text{CH}_2\text{---} \text{CH---} \text{O}$$

● HCl

RN 691872-66-9 CAPLUS

CN Piperazine, 1-[2-(4-fluorophenoxy)-2-phenylethyl]-4-[3-(trifluoromethyl)phenyl]-, hydrochloride (1:1) (CA INDEX NAME)

$$\begin{array}{c} \text{Ph} \\ \text{F}_3\text{C} \\ \text{N} \end{array} \begin{array}{c} \text{Ph} \\ \text{CH}_2\text{-CH-O} \end{array} \begin{array}{c} \text{F} \\ \text{CH}_3 \end{array}$$

● HCl

RN 866548-21-2 CAPLUS

CN Piperazine, 1-[3-(4-fluorophenoxy)-3-phenylpropyl]-4-phenyl- (CA INDEX NAME)

- 866548-22-3 CAPLUS RN
- Piperazine, 1-[3-(4-fluorophenoxy)-3-phenylpropyl]-4-phenyl-, CN hydrochloride (1:1) (CA INDEX NAME)

### ● HC1

- 866548-23-4 CAPLUS
  Piperazine, 1-phenyl-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]-, CN hydrochloride (1:1) (CA INDEX NAME)

#### HC1

- 866548-24-5 CAPLUS
- CN Piperazine, 1-[2-(4-fluorophenoxy)-2-phenylethyl]-4-phenyl- (CA INDEX NAME)

- RN 866548-25-6 CAPLUS
- Piperazine, 1-[2-(4-fluorophenoxy)-2-phenylethyl]-4-phenyl-, hydrochloride CN (1:1) (CA INDEX NAME)

HC1

RN 866548-26-7 CAPLUS

CN Piperazine, 1-(3-chlorophenyl)-4-[2-phenyl-2-[4-(trifluoromethyl)phenoxy]ethyl]- (CA INDEX NAME)

RN 866548-27-8 CAPLUS CN Piperazine, 1-(3-ch.

Piperazine, 1-(3-chlorophenyl)-4-[2-phenyl-2-[4-(trifluoromethyl)phenoxy]ethyl]-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

RN 866548-28-9 CAPLUS

CN Piperazine, 1-(2-methoxyphenyl)-4-[2-phenyl-2-[4-(trifluoromethyl)phenoxy]ethyl]- (CA INDEX NAME)

- RN 866548-29-0 CAPLUS
- CN Piperazine, 1-(2-methoxypheny1)-4-[2-pheny1-2-[4-(trifluoromethy1)phenoxy]ethy1]-, hydrochloride (1:1) (CA INDEX NAME)

### HC1

- RN 866548-30-3 CAPLUS
- CN Piperazine, 1-[2-phenyl-2-[4-(trifluoromethyl)phenoxy]ethyl]-4-[3-(trifluoromethyl)phenyl]- (CA INDEX NAME)

- RN 866548-31-4 CAPLUS
- CN Piperazine, 1-[2-pheny1-2-[4-(trifluoromethyl)phenoxy]ethyl]-4-[3-(trifluoromethyl)phenyl]-, hydrochloride (1:1) (CA INDEX NAME)

$$F_3C \longrightarrow N - CH_2 - CH - O - CF_3$$

### ● HCl

- RN 866548-36-9 CAPLUS
- CN Piperazine, 1-phenyl-4-[2-phenyl-2-[4-(trifluoromethyl)phenoxy]ethyl]-(CA INDEX NAME)

RN 866548-37-0 CAPLUS

CN Piperazine, 1-pheny1-4-[2-pheny1-2-[4-(trifluoromethy1)phenoxy]ethy1]-, hydrochloride (1:1) (CA INDEX NAME)

HC1

L3 ANSWER 9 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2004:344622 CAPLUS

DOCUMENT NUMBER: 140:357212

TITLE: Preparation of substituted anilinic piperidines as MCH

selective antagonists

INVENTOR(S): Marzabadi, Mohammad R.; Wetzel, John; Deleon, John E.;

Jiang, Yu; Chen, Chien-An; Lu, Kai

PATENT ASSIGNEE(S): Synaptic Pharmaceutical Corporation, USA

SOURCE:

U.S., 394 pp. CODEN: USXXAM

DOCUMENT TYPE:

Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3 PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
US 6727264 US 20040073036 US 20060041139 US 7105544 US 7067534	B1 A1 A9 B2 B1	20040427 20040415 20060223 20060912 20060627	US 2002-188434 US 2003-345063 US 2003-719358	20020703 20030114 20031121
US 20040186103 US 20060084649 US 7199135 US 20060217418	A1 A9 B2 A1	20040923 20060420 20070403 20060928	US 2004-753057 US 2005-541991	20040106
US 20070043080 PRIORITY APPLN. INFO.:	Al	20070222	US 2005-214968 US 2001-303091P US 2002-346997P	20050830 P 20010705 P 20020109
			WO 2002-US21063 US 2003-345063 US 2003-719358	A2 20020703 A2 20020703 A2 20030114 A1 20031121 W 20040106
OTHER SOURCE(S):	MARPAT	140:357212	WO 2004 00173	W 20040100

The title compds. [I (R1 = H, alkyl, aryl, etc.; R2 = alkyl, cyclopropyl; R3 = (un)substituted (hetero)aryl; A = H, F, Cl, Br, CN, etc.; X = O, NH; n = 0-5), II (W = III, IV (wherein R1 = H, Me, Et; X = O, NR3, CO, a bond; Y = H, (hetero)aryl; R3 = H, (hetero)aryl); R2 and A as above)] which are selective antagonists for melanin concentrating hormone-1 (MCH1) receptors.

were

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prepared Thus, reacting 2-methyl-N-[3-(4-piperidinyl)phenyl]propanamide (preparation given) with 4-chloro-3',4'-dimethylbutyrophenone in the presence of K2CO3 and NaI in DMF afforded 80% V which showed Ki of 3.9 nM in cloned rat MCH1 binding assay.

387826-65-5P 387826-66-6P 387826-67-7P 387826-68-8P 387826-69-9P 387826-73-5P 387826-74-6P 387826-79-1P 387826-80-4P 387826-81-5P 387826-82-6P 387826-85-9P 487049-74-1P 487049-80-9P 487049-81-0P

<sup>\*</sup> STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY - AVAILABLE VIA OFFLINE PRINT \*

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487049-83-2P 487049-84-3P 487049-85-4P
487049-86-5P 487049-87-6P 487049-88-7P
487049-89-8P 487049-90-1P 487049-91-2P
487049-92-3P 487049-95-6P 487049-99-0P
487050-01-1P 487050-07-7P 487050-08-8P
487050-09-9P 487050-20-4P 487050-22-6P
487050-28-2P 487050-33-9P 487050-34-0P
487050-35-1P 487050-36-2P 487050-39-5P
487051-81-0P 487051-83-2P 487051-85-4P
487052-31-3P 487056-49-5P 487057-25-0P
487057-26-1P 487057-32-9P 487057-33-0P
487057-35-2P 487057-36-3P 487057-37-4P
487057-38-5P 487057-40-9P 487057-41-0P
487057-45-4P 487057-47-6P 487057-49-8P
487057-50-1P 487057-51-2P 487057-52-3P
487057-53-4P 487057-55-6P 487057-56-7P
487057-57-8P 487057-59-0P 487057-60-3P
487057-62-5P 487057-65-8P 487057-66-9P
487057-67-0P 487057-68-1P 487057-70-5P
487057-71-6P 487057-72-7P 487057-73-8P
487057-76-1P 488098-61-9P 488098-62-0P
488098-63-1P 488098-64-2P 488098-65-3P
488098-67-5P 488098-69-7P 488098-70-0P
488098-71-1P 488098-72-2P 488098-73-3P
488098-74-4P 488098-76-6P 488098-77-7P
488098-78-8P 488098-79-9P 488098-81-3P
488098-82-4P
```

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of substituted anilinic piperidines as MCH selective antagonists)

RN 387826-65-5 CAPLUS

CN Propanamide, N-[3-[1-[(3S)-3-(3-chlorophenoxy)-3-phenylpropyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 387826-66-6 CAPLUS

CN Propanamide, N-[3-[1-[(3\$)-3-(4-chlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 387826-67-7 CAPLUS

CN Propanamide, 2-methyl-N-[3-[1-[(3S)-3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]-4-piperidinyl]phenyl]- (CA INDEX NAME)

## Absolute stereochemistry.

RN 387826-68-8 CAPLUS

CN Propanamide, N-[3-[1-[(3R)-3-(2,5-difluorophenoxy)-3-phenylpropyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

## Absolute stereochemistry.

RN 387826-69-9 CAPLUS

CN Propanamide, N-[3-[1-[(3R)-3-(3,4-dichlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-73-5 CAPLUS
- CN Propanamide, N-[3-]1-[(3S)-3-(4-fluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-74-6 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-(4-bromophenoxy)-3-phenylpropy1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 387826-79-1 CAPLUS

<12/04/2007>

Erich Leese

CN Propanamide, N-[3-[1-[(3R)-3-[2-fluoro-5-(trifluoromethyl)phenoxy]-3phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-80-4 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-[2-fluoro-5-(trifluoromethy1)phenoxy]-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-81-5 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-(2,5-difluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 387826-82-6 CAPLUS CN Propanamide, N-[3-[1-[(3R)-3-(3-chlorophenoxy)-3-phenylpropy1]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

## Absolute stereochemistry.

- 387826-85-9 CAPLUS RN
- CN Propanamide, N-[3-[1-[(3S)-3-(2-fluorophenoxy)-3-phenylpropyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

## Absolute stereochemistry.

- RN
- $\begin{array}{lll} 487049-74-1 & \text{CAPLUS} \\ \text{Propanamide, N-[3-[1-[3-(4-\text{chloropheny1})-3-(3,4-\text{difluorophenoxy})\text{propy1}]-4-} \end{array}$ CN piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487049-80-9 CAPLUS
  CN Propanamide, N-[3-[1-[3-(4-chlorophenoxy)-3-(4-fluoropheny1)propy1]-4piperidiny1)pleny1]-2-methy1- (CA INDEX NAME)
- RN 487049-81-0 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-fluorophenyl)-3-(2,3,4,5,6-pentafluorophenoxy)propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487049-83-2 CAPLUS CN Propagamide, N-13-11
- CN Propanamide, N-[3-[1-[3-(3,4-difluorophenoxy)-3-(4-fluorophenyl)propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487049-84-3 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-bromophenoxy)-3-(4-fluorophenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487049-85-4 CAPLUS
- CN Propanamide, N-[3-[1-[3-(3,4-dichlorophenoxy)-3-(4-fluoropheny1)propy1]-4-piperidiny1]pheny1]-2-methy1- (CA INDEX NAME)

- RN 487049-86-5 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-fluoropheny1)-3-[4-(trifluoromethy1)phenoxy]propy1]-4-piperidiny1]pheny1]-2-methy1- (CA INDEX NAME)

- RN 487049-87-6 CAPLUS
- CN Propanamide, N-[3-[1-[3-(3-bromophenoxy)-3-(4-fluorophenyl)propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487049-88-7 CAPLUS
CN Propanamide, N-[3-[1-[3-(4-fluorophenoxy)-3-(4-fluorophenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487049-89-8 CAPLUS
CN Propanamide, N-[3-[1-[3-(3-fluorophenoxy)-3-(4-fluorophenyl)propyl]-4piperidinvl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487049-90-1 CAPLUS
CN Propanamide, N-[3-[1-[3-(2,6-dichlorophenoxy)-3-(4-fluoropheny1)propy1]-4piperidiny1]pheny1]-2-methy1- (CA INDEX NAME)

RN 487049-91-2 CAPLUS

CN Propanamide, N-[3-[1-[3-(2,5-difluorophenoxy)-3-(4-fluorophenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487049-92-3 CAPLUS

CN Propanamide, N-[3-[1-[3-(3-chlorophenoxy)-3-(4-fluorophenyl)propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487049-95-6 CAPLUS

CN Propanamide, N-[3-[1-[3-(2,4-difluorophenoxy)-3-(4-fluorophenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487049-99-0 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-bromopheny1)-3-(2-chloro-4-methylphenoxy)propy1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487050-01-1 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-bromopheny1)-3-[4-(trifluoromethyl)phenoxy]propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

$$\begin{array}{c} \text{Br} \\ \text{0} \\ \text{0-CH-CH}_2\text{-CH}_2 \\ \text{N} \end{array}$$

- RN 487050-07-7 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-bromophenoxy)-3-(4-bromophenyl)propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487050-08-8 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-bromopheny1)-3-(4-chlorophenoxy)propy1]-4-piperidiny1]pheny1]-2-methy1- (CA INDEX NAME)

RN 487050-09-9 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-bromopheny1)-3-(4-fluorophenoxy)propy1]-4piperidiny1]pheny1]-2-methy1- (CA INDEX NAME)

RN 487050-20-4 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-bromophenoxy)-3-(4-chlorophenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487050-22-6 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-chlorophenoxy)-3-(4-chlorophenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487050-28-2 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-chlorophenoxy)-3-(4-methoxyphenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487050-33-9 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-bromophenoxy)-3-(4-methoxyphenyl)propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487050-34-0 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-methoxyphenyl)-3-[4-(trifluoromethyl)phenoxy]propyl]-4-piperidinyl]phenyl]-2-methyl-(NDEX NAME)

OMe

RN 487050-35-1 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-chloropheny1)-3-(4-fluorophenoxy)propy1]-4piperidiny1]pheny1]-2-methy1- (CA INDEX NAME)

RN 487050-36-2 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-fluorophenoxy)-3-(4-methoxyphenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487050-39-5 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-chlorophenyl)-3-[4-(trifluoromethyl)phenoxy]propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487051-81-0 CAPLUS
- CN Propanamide, N=[3=[1-[(2S)-2-(3-chlorophenoxy)-2-phenylethyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 487051-83-2 CAPLUS
- CN Propanamide, N-[3-[1-[(2R)-2-(4-fluorophenoxy)-2-phenylethyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 487051-85-4 CAPLUS
- CN Propanamide, N-[3-[1-[(2R)-2-(3-chlorophenoxy)-2-phenylethyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 487052-31-3 CAPLUS
- CN Propanamide, N-[3-[1-[4-(4-chlorophenoxy)-4-(4-chlorophenyl)butyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487056-49-5 CAPLUS

CN Propanamide, N-[3-[1-[(3S)-3-(3,4-dichlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

### Absolute stereochemistry.

RN 487057-25-0 CAPLUS

CN Propanamide, N-[3-[1-[6-(2-fluorophenoxy)-6-(2-fluorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-26-1 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-fluorophenoxy)-6-(2-fluorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-32-9 CAPLUS
- CN Propanamide, N-[3-[1-[6-(4-bromophenoxy)-6-phenylhexyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-33-0 CAPLUS
- CN Propanamide, N-[3-[1-[6-(4-fluorophenoxy)-6-(4-fluoropheny1)hexy1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-35-2 CAPLUS
- CN Propanamide, N-[3-[1-[6-(4-chlorophenoxy)-6-(4-chlorophenyl)hexyl]-4piperidinyl|phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-36-3 CAPLUS
- CN Propanamide, N-[3-[1-[6-(4-bromophenoxy)-6-(4-fluorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-37-4 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-chlorophenoxy)-6-(4-fluorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-38-5 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-chloropheny1)-6-(4-fluorophenoxy)hexy1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-40-9 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-chlorophenoxy)-6-phenylhexyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

i-Pr-C-NH

RN 487057-41-0 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-bromophenoxy)-6-(4-chlorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-45-4 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-fluorophenoxy)-6-phenylhexyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-47-6 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-fluorophenyl)-6-[2-fluoro-5-(trifluoromethyl)phenoxy]hexyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-49-8 CAPLUS CN Propagamide, N-13-1

CN Propanamide, N-[3-[1-[6-(2-fluorophenyl)-6-[2-fluoro-5-(trifluoromethyl)phenoxy]hexyl]-4-piperidinyl]phenyl]-2-methyl-(CA INDEX NAME)

RN 487057-50-1 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-chlorophenyl)-6-[2-fluoro-5-(trifluoromethyl)phenoxy]hexyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-51-2 CAPLUS

CN Propanamide, N-[3-[1-[6-(3-fluorophenyl)-6-[2-fluoro-5-(trifluoromethyl)phenoxy]hexyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-52-3 CAPLUS

CN Propanamide, N-[3-[1-[6-[2-fluoro-5-(trifluoromethyl)phenoxy]-6phenylhexyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-53-4 CAPLUS

CN Propanamide, N-[3-[1-[7-(2-fluorophenyl)-7-[2-fluoro-5-(trifluoromethyl)phenoxy]heptyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-55-6 CAPLUS
CN Propanamide, N-[3-[1-[5-(4-bromophenoxy)-5-(4-fluorophenyl)pentyl]-4piperiddinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-56-7 CAPLUS

CN Propanamide, N-[3-[1-[5-(4-chlorophenoxy)-5-(4-chlorophenyl)pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-57-8 CAPLUS
- CN Propanamide, N-[3-[1-[5-(4-chlorophenoxy)-5-phenylpenty1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-59-0 CAPLUS
- CN Propanamide, N-[3-[1-[5-(4-chloropheny1)-5-(4-fluorophenoxy)penty1]-4-piperidiny1]pheny1]-2-methy1- (CA INDEX NAME)

- RN 487057-60-3 CAPLUS
- CN Propanamide, N-[3-[1-[5-(4-bromophenoxy)-5-phenylpentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-62-5 CAPLUS
- CN Propanamide, N-[3-[1-[5-(4-fluorophenoxy)-5-(4-fluorophenyl)pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-65-8 CAPLUS

CN Propanamide, N-[3-[1-[5-[2-fluoro-5-(trifluoromethyl)phenoxy]-5-[4-(trifluoromethyl)phenyl]pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-66-9 CAPLUS
- CN Propanamide, N-[3-[1-[5-(3-chlorophenyl)-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-67-0 CAPLUS
- CN Propanamide, N-[3-[1-[5-(2-fluorophenyl)-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methyl-(CA INDEX NAME)

- RN 487057-68-1 CAPLUS
- CN Propanamide, N-[3-[1-[5-(3-fluorophenyl)-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-70-5 CAPLUS

CN Propanamide, N-[3-[1-[5-(4-chlorophenyl)-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-71-6 CAPLUS

CN Propanamide, N-[3-[1-[5-(4-fluorophenyl)-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-72-7 CAPLUS

CN Propanamide, N-[3-[1-[5-(4-bromophenoxy)-5-(4-chlorophenyl)pentyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-73-8 CAPLUS

CN Propanamide, N-[3-[1-[5-(4-chlorophenoxy)-5-(4-fluorophenyl)pentyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-76-1 CAPLUS

CN Propanamide, N-[3-[1-[5-(4-fluorophenoxy)-5-phenylpenty1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

$$\begin{array}{c} \text{Ph} \\ \text{i-Pr-C-NH} \end{array}$$

RN 488098-61-9 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-fluorophenoxy)-6-(2-fluorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

HC1

RN 488098-62-0 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-fluorophenoxy)-6-(4-fluorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

HC1

RN 488098-63-1 CAPLUS CN Propanamide, N-[3-[1-[6-(4-chlorophenoxy)-6-(4-fluorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

HC1

- RN 488098-64-2 CAPLUS
- CN Propanamide, N-[3-[1-[6-(4-chlorophenyl)-6-(4-fluorophenoxy)hexyl]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

HC1

RN 488098-65-3 CAPLUS

<12/04/2007>

CN Propanamide, N-[3-[1-[6-(4-chlorophenoxy)-6-phenylhexy1]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

$$\begin{array}{c} \text{Ph} \\ \text{O} \\ \text{i-Pr-C-NH} \end{array}$$

● HCl

RN 488098-67-5 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-fluorophenoxy)-6-phenylhexyl]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

RN 488098-69-7 CAPLUS

CN Propanamide, N-[3-[1-[6-(2-fluorophenyl)-6-[2-fluoro-5-(trifluoromethyl)phenoxy]hexyl]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

HC1

RN 488098-70-0 CAPLUS

CN Propanamide, N-[3-[1-[5-(4-bromophenoxy)-5-(4-fluorophenyl)pentyl]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

10/513699

HC1

- RN 488098-71-1 CAPLUS
  CN Propanamide, N-[3-[1-[5-(4-chlorophenoxy)-5-(4-chlorophenyl)pentyl]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)
- C1 O-CH-(CH<sub>2</sub>)<sub>4</sub>-N NH-C-Pr-

HCl

RN 488098-72-2 CAPLUS

- CN Propanamide, N-[3-[1-[5-(4-chlorophenoxy)-5-phenylpentyl]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)
- Ph C1

HC1

- RN 488098-73-3 CAPLUS
- CN Propanamide, N-[3-[1-[5-(4-chloropheny1)-5-(4-fluorophenoxy)penty1]-4-piperidiny1]pheny1]-2-methy1-, hydrochloride (1:1) (CA INDEX NAME)

### HC1

- 488098-74-4 CAPLUS RN
- Propanamide, N-[3-[1-[5-(4-bromophenoxy)-5-phenylpentyl]-4-CN piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

# ● HCl

- RN 488098-76-6 CAPLUS
- CN Propanamide, N-[3-[1-[5-(4-fluorophenoxy)-5-(4-fluorophenyl)pentyl]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

### HC1

- RN
- 488098-77-7 CAPLUS Propanamide, N-[3-[1-[5-(3-chloropheny1)-5-[2-fluoro-5-CN (trifluoromethy1)phenoxy]penty1]-4-piperidiny1]pheny1]-2-methy1-, hydrochloride (1:1) (CA INDEX NAME)

10/513699

● HC1

RN 488098-78-8 CAPLUS

CN Propanamide, N-[3-[1-[5-(2-fluorophenyl)-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methylhydrochloride (1:1) (CA INDEX NAME)

● HC1

RN 488098-79-9 CAPLUS

CN Propanamide, N-[3-[1-[5-(3-fluoropheny])-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

HC1.

RN 488098-81-3 CAPLUS

<12/04/2007>

Erich Leese

CN Propanamide, N-[3-[1-[5-(4-chloropheny])-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

• HCl

RN 488098-82-4 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-fluoropheny1)-6-[2-fluoro-5-(trifluoromethyl)]phenoxy]hexy]]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

HCl

REFERENCE COUNT:

25 THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

PUBLISHER:

```
ANSWER 10 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN
ACCESSION NUMBER:
                        2004:170822 CAPLUS
DOCUMENT NUMBER:
                        140:417233
TITLE:
                        Synthesis and biological evaluation of
```

2-(4-fluorophenoxy)-2-phenyl-ethyl piperazines as serotonin-selective reuptake inhibitors with a potentially improved adverse reaction profile

Dorsey, James M.; Miranda, Maria G.; Cozzi, Nicholas AUTHOR(S):

V.; Pinney, Kevin G.

CORPORATE SOURCE: Department of Chemistry and Biochemistry and The Center for Drug Discovery, Baylor University, Waco,

TX, 76798-7348, USA SOURCE:

Bioorganic & Medicinal Chemistry (2004), 12(6), 1483-1491

Elsevier Ltd.

CODEN: BMECEP; ISSN: 0968-0896

DOCUMENT TYPE: Journal LANGUAGE: English

OTHER SOURCE(S): CASREACT 140:417233

Three new 2-(4-fluorophenoxy)-2-phenyl-Et piperazines, 1-(3-chlorophenv1)-4-[2-(4-fluorophenoxy)-2-phenvlethv1]-piperazine, 1-(2-(4-fluorophenoxy)-2-phenylethyl)-4-(2-methoxyphenyl)-piperazine, and 1-[2-(4-fluorophenoxy)-2-phenylethyl]-4-(3-trifluoromethylphenyl)piperazine, modeled after the potent antidepressant fluoxetine and coupled with several functionalized piperazines, have been prepared by chemical synthesis as selective serotonin reuptake inhibitors (SSRIs) with a potentially improved adverse reaction profile. Typical SSRIs, although very effective in the treatment of depression, still face the troublesome side effect of sexual dysfunction. A number of pharmacol. agents-notably, drugs in the piperazine class-have been used to reverse SSRI-induced sexual dysfunction, and evidence for developing an improved SSRI by coupling a fluoxetine congener with the pharmacophore of a reversal agent holds promise. Preliminary data indicates that the hydrochloride (HCl) salts of piperazines exhibit single-site binding at the site of the serotonin reuptake transporter (SERT). However, each of the three compds. are much less potent than typical SSRIs, showing micromolar (µM) affinity for the SERT with IC50 values of 1.45 µM, 3.27 µM, and 9.56 μM, resp. Further biol. evaluation of piperazine compds. is needed before definitive conclusions can be made with regard to each compound's potential for use as an SSRI-type candidate which is devoid of sexual side effects. Nevertheless, the initial findings are quite encouraging, thus lending credence to the idea of hybridizing an SSRI congener with that of

sexual dysfunction. 691872-62-5P 691872-64-7P 691872-66-9P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

the pharmacophore of an agent known to reverse or treat SSRI-induced

(synthesis and structure-activity relationship of

2-(4-fluorophenoxy)-2-Ph-Et piperazines as serotonin-selective reuptake inhibitors with a potentially improved adverse reaction profile) 691872-62-5 CAPLUS

RN

Piperazine, 1-(3-chlorophenyl)-4-(2-(4-fluorophenoxy)-2-phenylethyl]-, CN hydrochloride (1:1) (CA INDEX NAME)

CN

# ● HCl

RN 691872-64-7 CAPLUS

Piperazine, 1-[2-(4-fluorophenoxy)-2-phenylethyl]-4-(2-methoxyphenyl)-, hydrochloride (1:1) (CA INDEX NAME)

# ● HCl

RN 691872-66-9 CAPLUS

CN Piperazine, 1-[2-(4-fluorophenoxy)-2-phenylethyl]-4-[3-(trifluoromethyl)phenyl]-, hydrochloride (1:1) (CA INDEX NAME)

$$F_3C \\ \hline \\ N \\ \hline \\ CH_2 \\ \hline \\ CH \\ -O \\ \hline \\ CH \\ -O$$

# ● HCl

IT 691872-56-7P 691872-58-9P 691872-60-3P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(synthesis and structure-activity relationship of 2-(4-fluorophenoxy)-2-Ph-Et piperazines as serotonin-selective reuptake inhibitors with a potentially improved adverse reaction profile)

RN 691872-56-7 CAPLUS

<12/04/2007>

CN Piperazine, 1-(3-chlorophenyl)-4-[2-(4-fluorophenoxy)-2-phenylethyl]- (CA INDEX NAME)

RN 691872-58-9 CAPLUS

CN Piperazine, 1-[2-(4-fluorophenoxy)-2-phenylethyl]-4-(2-methoxyphenyl)-(CA INDEX NAME)

$$\stackrel{\text{OMe}}{\underset{\text{N}}{\longrightarrow}} \text{N-CH}_2 - \text{CH-O}$$

RN 691872-60-3 CAPLUS

CN Piperazine, 1-[2-(4-fluorophenoxy)-2-phenylethyl]-4-[3-(trifluoromethyl)phenyl]- (CA INDEX NAME)

$$\begin{array}{c|c} & \text{Ph} & \\ & \text{F3C} & \\ & \text{N} & \text{CH}_2\text{-CH-O} \end{array}$$

REFERENCE COUNT:

14 THERE ARE 14 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 11 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2004:162444 CAPLUS

DOCUMENT NUMBER: 140:212060

TITLE: DNA encoding a human melanin concentrating hormone receptor (MCH1) and uses thereof and preparation of 4-phenylpiperidine derivatives as human MCH1 receptor

antagonists

INVENTOR(S): Salon, John A.; Laz, Thomas M.; Nagorny, Raisa;

Wilson, Amy E.; Craig, Douglas A. PATENT ASSIGNEE(S):

SOURCE: U.S. Pat. Appl. Publ., 180 pp., Cont.-in-part of U.S.

Ser. No. 899,732. CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: FAMILY ACC. NUM. COUNT: PATENT INFORMATION:					English 4															
PATENT NO.										APPL		DATE								
US WO	20040038855				A1 20040226				US 2003-341751 WO 1999-US31169						20030114					
	W:	CZ, IS, MG,	DE, JP, MK,	DK, KE, MN,	EE, KG, MW,	ES, KP, MX,	AZ, FI, KR, NO, TZ,	GB, KZ, NZ,	GD, LC, PL,	GE, LK, PT,	GH, LR, RO,	GM, LS, RU,	HR, LT, SD,	HU, LU, SE,	ID, LV, SG,	IL, MA,	IN, MD,			
	RW:	GH, DK,	GM, ES,	KE, FI, CM,	LS, FR, GA,	MW, GB,	SD, GR, GW,	SL, IE,	SZ, IT,	TZ, LU,	UG, MC,	ZW, NL,	AT, PT,	BE,	CH,					
WO					A2 20040805					US 2001-899732 WO 2004-US724						20010705 20040114				
		CN, GE, LK, NO, TJ, AT, IT,	CO, GH, LR, NZ, TM, BE, LU,	CR, GM, LS, OM, TN, BG, MC,	CU, HR, LT, PG, TR, CH, NL,	CZ, HU, LU, PH, TT, CY, PT,	AU, DE, ID, LV, PL, TZ, CZ, RO,	DK, IL, MA, PT, UA, DE, SE,	DM, IN, MD, RO, UG, DK, SI,	DZ, IS, MG, RU, US, EE, SK,	EC, JP, MK, SC, UZ, ES, TR,	EE, KE, MN, SD, VC, FI, BF,	EG, KG, MW, SE, VN, FR, BJ,	ES, KP, MX, SG, YU, GB, CF,	FI, KR, MZ, SK, ZA, GR, CG,	GB, KZ, NA, SL, ZM, HU, CI,	GD, LC, NI, SY, ZW IE, CM,			
PRIORIT		GW, ML, MR, NE, S SZ, TZ, UG, ZM, Z				W, AM, AZ, BY, KG,														

GI

AB

receptor, a purified human MCH1 receptor, vectors comprising isolated nucleic acid encoding a human MCH1 receptor, cells comprising such vectors, antibodies directed to a human MCH1 receptor, nucleic acid probes useful for detecting nucleic acid encoding human MCH1 receptors, antisense oligonucleotides complementary to unique sequences of nucleic acid encoding human MCH1 receptors, transgenic, nonhuman animals which express DNA encoding a normal or mutant human MCH1 receptor, methods of isolating a human MCH1 receptor, methods of treating an abnormality that is linked to the activity of a human MCH1 receptor, as well as methods of determining binding of compds. to mammalian MCH1 receptors. This invention further provides a method of treating a subject suffering from urinary incontinence which comprises administering to the subject an amount of an MCH1 antagonist effective to treat the subject's urinary incontinence or overactive bladder. Various 4-phenylpiperidine derivs., e.g (I), were synthesized and tested as human MCH1 receptor antagonists. IΤ 387826-65-5P, N-[3-[1-[(3S)-3-(3-Chlorophenoxy)-3-phenylpropyl]-4piperidinyl]phenyl]-2-methylpropanamide 387826-66-6P, N-[3-[1-[(3S)-3-(4-Chlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2methylpropanamide 387826-67-7P, 2-Methyl-N-[3-[1-[(3S)-3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]-4piperidinvl|phenvl|propanamide 387826-68-8P, N-(3-(1-(3R)-3-(2.5-Difluorophenoxy)-3-phenylpropyl]-4piperidinyllphenyll-2-methylpropanamide 387826-69-9P. N-[3-[1-[(3R)-3-(3,4-Dichlorophenoxy)-3-phenylpropy1]-4piperidinyl]phenyl]-2-methylpropanamide 387826-73-5P, N-[3-[1-[(3S)-3-(4-Fluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2methylpropanamide 387826-74-6P, N-(3-(1-(3S)-3-(4-Bromophenoxy)-3-phenylpropyl)-4-piperidinylphenyl]-2methylpropanamide 387826-79-1P, N-[3-[1-[(3R)-3-[2-Fluoro-5-(trifluoromethyl)phenoxy]-3-phenylpropyl]-4piperidinyl]phenyl]-2-methylpropanamide 387826-80-4P, N-[3-[1-[(3S)-3-[2-Fluoro-5-(trifluoromethyl)phenoxy]-3-phenylpropyl]-4piperidinvl]phenvl]-2-methvlpropanamide 387826-81-5P,

N-[3-[1-[(3R)-3-(3-Chlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-

N-[3-[1-[(3S)-3-(2-Fluorophenoxy)-3-phenylpropy1]-4-piperidiny1]pheny1]-2-

This invention provides an isolated nucleic acid encoding a human MCH1

<12/04/2007> Erich Leese

methylpropanamide 387826-85-9P,

methylpropanamide

N-[3-[1-[(38)-3-(2,5-Difluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methylpropanamide 387826-82-6P,

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(DNA encoding human melanin concentrating hormone receptor (MCH1) and uses thereof and preparation of phenylpiperidine derivs. as human MCH1 antagonists)

RN 387826-65-5 CAPLUS

CN Propanamide, N-[3-[1-[(3S)-3-(3-chlorophenoxy)-3-phenylpropyl]-4piperidinyl|phenyl|-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 387826-66-6 CAPLUS

CN Propanamide, N-[3-[1-[(3S)-3-(4-chlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 387826-67-7 CAPLUS

CN Propanamide, 2-methyl-N-[3-[1-[(3S)-3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]-4-piperidinyl]phenyl]- (CA INDEX NAME)

Absolute stereochemistry.

10/513699

RN 387826-68-8 CAPLUS
CN Propanamide, N-[3-[1-[(3R)-3-(2,5-difluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl-2-methyl- (CA INDEX NAME)

### Absolute stereochemistry.

- RN 387826-69-9 CAPLUS
- CN Propanamide, N-[3-[1-[(3R)-3-(3,4-dichlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

### Absolute stereochemistry.

RN 387826-73-5 CAPLUS

<12/04/2007>

Erich Leese

CN Propanamide, N-[3-[1-[(3\$)-3-(4-fluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-74-6 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-(4-bromophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-79-1 CAPLUS
- CN Propanamide, N-[3-[1-[(3R)-3-[2-fluoro-5-(trifluoromethyl)phenoxy]-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-80-4 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-[2-fluoro-5-(trifluoromethyl)phenoxy]-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

### Absolute stereochemistry.

- RN 387826-81-5 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-(2,5-difluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

# Absolute stereochemistry.

- RN 387826-82-6 CAPLUS
- CN Propanamide, N-[3-[1-[(3R)-3-(3-chlorophenoxy)-3-phenylpropy1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

## Absolute stereochemistry.

RN 387826-85-9 CAPLUS

CN Propanamide, N-[3-]1-[(3S)-3-(2-fluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

L3 ANSWER 12 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2003:334519 CAPLUS

DOCUMENT NUMBER . 138 - 298124

TITLE: Human melanin concentrating hormone receptor MCH1, its

DNA, its synthetic ligands and diagnostic and

therapeutic uses thereof

Borowsky, Beth; Blackburn, Thomas P.; Ogozalek, INVENTOR(S):

Kristine

PATENT ASSIGNEE(S): USA

SOURCE: U.S. Pat. Appl. Publ., 193 pp., Cont.-in-part of U.S.

Ser. No. 610,635. CODEN: USXXCO DOCUMENT TYPE: Patent

LANGUAGE: English FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

PATE	KIND		DATE			APPLICATION NO.					DATE								
US 6 WO 2	US 20030082623 US 6221613 WO 2000039279 WO 2000039279					A1 B1 A2 A3		20030501 20010424 20000706 20001102		US 2001-899732 US 1998-224426 WO 1999-US31169					20010705 19981231 19991230				
	W:	AE, CZ, IS, MG, SL, GH,	AL, DE, JP, MK, TJ, GM,	DK, KE, MN, TM, KE,	AT, EE, KG, MW, TR, LS,	AU, ES, KP, MX, TT,	AZ, FI, KR, NO, TZ, SD, GR,	BA, GB, KZ, NZ, UA, SL,	GD, LC, PL, UG, SZ,	GE, LK, PT, US, TZ,	GH, LR, RO, UZ, UG,	GM, LS, RU, VN, ZW,	HR, LT, SD, YU, AT,	HU, LU, SE, ZA, BE,	ID, LV, SG, ZW CH,	IL, MA, SI,	IN, MD, SK, DE,		
US 2								ML, 0424	MR, NE, SN, TD, TG US 2001-29314 US 2003-341751 US 1998-224426 WO 1999-US31169 US 2000-610635 US 2001-899732					1	20011220 20030114 A2 19981231 A2 19991230 A2 20000705 A1 20010705				

This invention provides an isolated nucleic acid encoding a human MCH1 receptor, a purified human MCH1 receptor, vectors comprising isolated nucleic acid encoding a human MCH1 receptor, cells comprising such vectors, antibodies directed to a human MCH1 receptor, nucleic acid probes useful for detecting nucleic acid encoding human MCH1 receptors, antisense oligonucleotides complementary to unique sequences of nucleic acid encoding human MCH1 receptors, transgenic, nonhuman animals which express DNA encoding a normal or mutant human MCH1 receptor, methods of isolating a human MCH1 receptor, methods of treating an abnormality that is linked to the activity of a human MCH1 receptor, as well as methods of determining binding of compds. to mammalian MCH1 receptors. This invention provides a method of modifying the feeding behavior of a subject which comprises administering to the subject an amount of an MCH1 antagonist effective to decrease the body mass of the subject and/or decrease the consumption of food by the subject. This invention further provides a method of treating a subject suffering from depression and/or anxiety which comprises administering to the subject an amount of an MCH1 antagonist effective to treat the subject's depression and/or anxiety.

<sup>387826-65-5</sup>P 387826-66-6P 387826-67-7P 387826-68-8P 387826-69-9P 387826-73-5P

387826-74-6P 387826-79-1P 387826-80-4P 387826-81-5P 387826-82-6P 387826-85-9P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(Uses)
(Numan melanin concentrating hormone receptor MCH1, its DNA, its synthetic ligands and diagnostic and therapeutic uses thereof)

RN 387826-65-5 CAPLUS

CN Propanamide, N-[3-[1-[(3S)-3-(3-chlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 387826-66-6 CAPLUS

CN Propanamide, N-[3-[1-[(3S)-3-(4-chlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 387826-67-7 CAPLUS

CN Propanamide, 2-methyl-N-[3-[1-[(3S)-3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]-4-piperidinyl]phenyl]- (CA INDEX NAME)

Absolute stereochemistry.

10/513699

RN 387826-68-8 CAPLUS
CN Propanamide, N-[3-[1-[(3R)-3-(2,5-difluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl-2-methyl- (CA INDEX NAME)

#### Absolute stereochemistry.

- RN 387826-69-9 CAPLUS
- CN Propanamide, N-[3-[1-[(3R)-3-(3,4-dichlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

### Absolute stereochemistry.

RN 387826-73-5 CAPLUS

<12/04/2007>

Erich Leese

CN Propanamide, N-[3-[1-[(3\$)-3-(4-fluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-74-6 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-(4-bromophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-79-1 CAPLUS
- CN Propanamide, N-[3-[1-[(3R)-3-[2-fluoro-5-(trifluoromethyl)phenoxy]-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-80-4 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-[2-fluoro-5-(trifluoromethyl)phenoxy]-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

#### Absolute stereochemistry.

- RN 387826-81-5 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-(2,5-difluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

## Absolute stereochemistry.

- RN 387826-82-6 CAPLUS
- CN Propanamide, N-[3-[1-[(3R)-3-(3-chlorophenoxy)-3-phenylpropy1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

### Absolute stereochemistry.

RN 387826-85-9 CAPLUS

CN Propanamide, N-[3-]1-[(3S)-3-(2-fluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

L3 ANSWER 13 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2003:42108 CAPLUS

DOCUMENT NUMBER: 138:106601

TITLE: Preparation of substituted anilinic piperidines as MCH

selective antagonists

INVENTOR(S): Marzabadi, Mohammad R.; Wetzel, John; Deleon, John E.;

Jiang, Yu

Synaptic Pharmaceutical Corporation, USA PATENT ASSIGNEE(S):

SOURCE: PCT Int. Appl., 771 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 3

PATENT INFORMATION:

PATENT NO.	KIND DATE	APPLICATION NO.	DATE
CO, CR, CU, GM, HR, HU, LS, LT, LU, PL, PT, RO, UA, UG, UZ, RW: GH, GM, KE,	A1 20030116 AM, AT, AU, AZ, CZ, DE, DK, DM, ID, IL, IN, IS, LV, MA, MD, MG, RU, SD, SE, SG, VN, YU, ZA, ZM, LS, MW, MZ, SD,	SL, SZ, TZ, UG, ZM, ZW,	GD, GE, GH, LC, LK, LR, NZ, OM, PH, TR, TT, TZ, AT, BE, BG,
	TR, BF, BJ, CF,	FI, FR, GB, GR, IE, IT, CG, CI, CM, GA, GN, GQ,	GW, ML, MR,
AU 2002316531	B2 20070913	AU 2002-316531	20020703 20020703
R: AT, BE, CH, IE, SI, LT,	LV, FI, RO, MK,	EP 2002-746843 GB, GR, IT, LI, LU, NL, CY, AL, TR, BG, CZ, EE, BR 2002-10869	SE, MC, PT, SK
BR 2002010869 JP 2004536104 HU 2004001880	T 20041202	TD 2002 E10020	20020703 20020703 20020703
CN 1671386 NZ 530221 US 20040073036 US 20060041139	A 20050921 A 20060331 A1 20040415 A9 20060223	HU 2003-310038 EA 2003-9860 CN 2002-817212 NZ 2002-530221 US 2003-345063	20020703 20020703 20020703 20030114
NO 2004000028 US 20040186103 US 20060084649	A 20050307 A 20040304 A1 20040923 A9 20060420	NO 2004-28 US 2004-753057	20031218
US 7199135 IN 2004CN00230 PRIORITY APPLN. INFO.:	A 20051209	US 2001-899794 US 2002-42582 US 2001-303091P	A 20020109 P 20010705
OTHER SOURCE(S):	MARPAT 138:1066	US 2002-346997P US 2002-188434 WO 2002-US21063 US 2003-345063	W 20020703

G1

AB The title compds. [I (R1 = H, alkyl, aryl, etc.; R2 = alkyl, cyclopropyl; R3 = (un)substituted (hetero)aryl; A = H, F, Cl, Br, CN, etc.; X = O, NH; n = 0-5), II (R1 = (un)substituted (hetero)aryl; R2, A, n as above), etc.] which are selective antagonists for melanin concentrating hormone-1 (MCH1)

receptors, were prepared and formulated. Thus, reacting 2-methyl-M-[3-(4-piperidinyl)phenyl]propanamide (preparation given) with 4-chloro-3',4'-dimethylbutyrophenone in the presence of K2CO3 and NaI in

DMF afforded 80% II [R1 = R1 = 3,4-Me2C6H3; R2 = iso-Pr; A = H; n = 2]

which showed Ki of 3.9 nM in cloned rat MCH1 binding assay.

387826-65-5P 387826-66-6P 387826-67-7P 387826-68-8P 387826-69-9P 387826-73-5P 387826-74-6P 387826-79-1P 387826-80-4P 387826-81-5P 387826-82-6P 387826-85-9P 487049-74-1P 487049-80-9P 487049-81-0P 487049-83-2P 487049-84-3P 487049-85-4P 487049-86-5P 487049-87-6P 487049-88-7P 487049-89-8P 487049-90-1P 487049-91-2P 487049-92-3P 487049-95-6P 487049-99-0P 487050-01-1P 487050-07-7P 487050-08-8P 487050-09-9P 487050-20-4P 487050-22-6P 487050-28-2P 487050-33-9P 487050-34-0P 487050-35-1P 487050-36-2P 487050-39-5P 487051-81-0P 487051-83-2P 487051-85-4P 487052-31-3P 487056-49-5P 487057-25-0P 487057-26-1P 487057-32-9P 487057-33-0P 487057-35-2P 487057-36-3P 487057-37-4P 487057-38-5P 487057-40-9P 487057-41-0P

#### 10/513699

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487057-45-4P 487057-47-6P 487057-49-8P 487057-50-1P 487057-55-1P 487057-55-3P 487057-55-3P 487057-55-3P 487057-55-3P 487057-55-3P 487057-55-6P 487057-56-6P 487057-66-3P 487057-66-5P 487057-66-5P 487057-66-5P 487057-66-1P 487057-70-5P 487057-71-6P 487057-71-2P 487057-71-6P 487057-71-2P 487057-71-6P 487057-71-2P 487057-71-6P 487057-71-2P 487057-71-3P 487057-71-3P 487057-71-3P 487057-71-3P 487057-71-3P 487057-71-3P 487057-71-3P 487057-71-3P 488098-61-9P 488098-65-3P 488098-71-3P 48809
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RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of substituted anilinic piperidines as MCH selective antagonists)

RN 387826-65-5 CAPLUS

CN Propanamide, N-[3-[1-[(3S)-3-(3-chlorophenoxy)-3-phenylpropyl]-4-piperidinylphenyll-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-66-6 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-(4-chlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Erich Leese

Absolute stereochemistry.

RN 387826-67-7 CAPLUS

<12/04/2007>

CN Propanamide, 2-methyl-N-[3-[1-[(3S)-3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]-4-piperidinyl]phenyl]- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-68-8 CAPLUS
- CN Propanamide, N-[3-[1-[(3R)-3-(2,5-difluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-69-9 CAPLUS
- CN Propanamide, N-[3-[1-[(3R)-3-(3,4-dichlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 387826-73-5 CAPLUS

CN Propanamide, N-[3-[1-[(3S)-3-(4-fluorophenoxy)-3-phenylpropy1]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

## Absolute stereochemistry.

- 387826-74-6 CAPLUS RN
- Propanamide, N-[3-[1-[(3S)-3-(4-bromophenoxy)-3-phenylpropy1]-4-bromophenoxy]CN piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

#### Absolute stereochemistry.

- RN
- $387826-79-1 \quad CAPLUS \\ Propanamide, \ N-[3-[1-[(3R)-3-[2-fluoro-5-(trifluoromethyl)phenoxy]-3-[3-fluoro-5-(trifluoromethyl)p$ CN phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-80-4 CAPLUS
- CN Propanamide, N=[3=[1-[(3S)-3-[2-fluoro-5-(trifluoromethy1)phenoxy]-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-81-5 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-(2,5-difluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 387826-82-6 CAPLUS CN Propanamide, N-[3-[1-[(3R)-3-(3-chlorophenoxy)-3-phenylpropy1]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

## Absolute stereochemistry.

- 387826-85-9 CAPLUS RN
- CN Propanamide, N-[3-[1-[(3S)-3-(2-fluorophenoxy)-3-phenylpropyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

#### Absolute stereochemistry.

- RN
- $\begin{array}{lll} 487049-74-1 & \text{CAPLUS} \\ \text{Propanamide, N-[3-[1-[3-(4-\text{chloropheny1})-3-(3,4-\text{difluorophenoxy})\text{propy1}]-4-} \end{array}$ CN piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487049-80-9 CAPLUS
  CN Propanamide, N-[3-[1-[3-(4-chlorophenoxy)-3-(4-fluoropheny1)propy1]-4piperidiny1)pleny1]-2-methy1- (CA INDEX NAME)
- RN 487049-81-0 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-fluoropheny1)-3-(2,3,4,5,6pentafluorophenoxy)propy1]-4-piperidiny1]pheny1]-2-methy1-NAME)

- RN 487049-83-2 CAPLUS
- CN Propanamide, N-[3-[1-[3-(3,4-difluorophenoxy)-3-(4-fluorophenyl)propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487049-84-3 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-bromophenoxy)-3-(4-fluorophenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487049-85-4 CAPLUS
- CN Propanamide, N-[3-[1-[3-(3,4-dichlorophenoxy)-3-(4-fluorophenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487049-86-5 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-fluorophenyl)-3-[4-(trifluoromethyl)phenoxy]propyl]-4-piperidinyl]phenyl]-2-methyl-(NDEX NAME)

- RN 487049-87-6 CAPLUS
- CN Propanamide, N-[3-[1-[3-(3-bromophenoxy)-3-(4-fluorophenyl)propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487049-88-7 CAPLUS
CN Propanamide, N-[3-[1-[3-(4-fluorophenoxy)-3-(4-fluorophenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487049-89-8 CAPLUS
CN Propanamide, N-[3-[1-[3-(3-fluorophenoxy)-3-(4-fluorophenyl)propyl]-4piperidinvl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487049-90-1 CAPLUS CN Propagamide, N-13-11

Propanamide, N-[3-[1-[3-(2,6-dichlorophenoxy)-3-(4-fluorophenyl)propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487049-91-2 CAPLUS

CN Propanamide, N-[3-[1-[3-(2,5-difluorophenoxy)-3-(4-fluorophenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487049-92-3 CAPLUS

CN Propanamide, N-[3-[1-[3-(3-chlorophenoxy)-3-(4-fluorophenyl)propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487049-95-6 CAPLUS

CN Propanamide, N-[3-[1-[3-(2,4-difluorophenoxy)-3-(4-fluorophenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

$$\begin{array}{c} F \\ O \\ O \\ CH \\ CH_2 \\ CH_2 \\ - NH \\ \end{array}$$

- RN 487049-99-0 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-bromopheny1)-3-(2-chloro-4-methylphenoxy)propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487050-01-1 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-bromopheny1)-3-[4-(trifluoromethyl)phenoxy[propy1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487050-07-7 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-bromophenoxy)-3-(4-bromopheny1)propy1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487050-08-8 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-bromopheny1)-3-(4-chlorophenoxy)propy1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487050-09-9 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-bromopheny1)-3-(4-fluorophenoxy)propy1]-4piperidiny1]pheny1]-2-methy1- (CA INDEX NAME)

RN 487050-20-4 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-bromophenoxy)-3-(4-chlorophenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487050-22-6 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-chlorophenoxy)-3-(4-chlorophenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487050-28-2 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-chlorophenoxy)-3-(4-methoxyphenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487050-33-9 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-bromophenoxy)-3-(4-methoxyphenyl)propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487050-34-0 CAPLUS
- CN Propanamide, N-[3-[1-[3-(4-methoxyphenyl)-3-[4-(trifluoromethyl)phenoxy]propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

OMe

RN 487050-35-1 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-chloropheny1)-3-(4-fluorophenoxy)propy1]-4piperidiny1]pheny1]-2-methy1- (CA INDEX NAME)

RN 487050-36-2 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-fluorophenoxy)-3-(4-methoxyphenyl)propyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487050-39-5 CAPLUS

CN Propanamide, N-[3-[1-[3-(4-chlorophenyl)-3-[4-(trifluoromethyl)phenoxy]propyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487051-81-0 CAPLUS
- CN Propanamide, N-[3-[1-[(2S)-2-(3-chlorophenoxy)-2-phenylethyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 487051-83-2 CAPLUS
- CN Propanamide, N-[3-[1-[(2R)-2-(4-fluorophenoxy)-2-phenylethyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 487051-85-4 CAPLUS
- CN Propanamide, N-[3-[1-[(2R)-2-(3-chlorophenoxy)-2-phenylethyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 487052-31-3 CAPLUS
- CN Propanamide, N-[3-[1-[4-(4-chlorophenoxy)-4-(4-chlorophenyl)butyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487056-49-5 CAPLUS

CN Propanamide, N-[3-[1-[(3S)-3-(3,4-dichlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

### Absolute stereochemistry.

- RN 487057-25-0 CAPLUS
- CN Propanamide, N-[3-[1-[6-(2-fluorophenoxy)-6-(2-fluorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-26-1 CAPLUS
- CN Propanamide, N-[3-[1-[6-(4-fluorophenoxy)-6-(2-fluorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-32-9 CAPLUS
- CN Propanamide, N-[3-[1-[6-(4-bromophenoxy)-6-phenylhexy1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-33-0 CAPLUS
- CN Propanamide, N-[3-[1-[6-(4-fluorophenoxy)-6-(4-fluorophenyl)hexyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-35-2 CAPLUS
- CN Propanamide, N-[3-[1-[6-(4-chlorophenoxy)-6-(4-chlorophenyl)hexyl]-4piperidinyl|phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-36-3 CAPLUS
- CN Propanamide, N-[3-[1-[6-(4-bromophenoxy)-6-(4-fluorophenyl)hexyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-37-4 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-chlorophenoxy)-6-(4-fluorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-38-5 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-chloropheny1)-6-(4-fluorophenoxy)hexy1]-4piperidiny1]pheny1]-2-methy1- (CA INDEX NAME)

RN 487057-40-9 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-chlorophenoxy)-6-phenylhexyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

i-Pr-C-NH

RN 487057-41-0 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-bromophenoxy)-6-(4-chlorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-45-4 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-fluorophenoxy)-6-phenylhexyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-47-6 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-fluorophenyl)-6-[2-fluoro-5-(trifluoromethyl)phenoxy]hexyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-49-8 CAPLUS CN Propagamide, N-13-1

CN Propanamide, N-[3-[1-[6-(2-fluorophenyl)-6-[2-fluoro-5-(trifluoromethyl)phenoxy]hexyl]-4-piperidinyl]phenyl]-2-methyl-(CA INDEX NAME)

RN 487057-50-1 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-chlorophenyl)-6-[2-fluoro-5-(trifluoromethyl)phenoxy]hexyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-51-2 CAPLUS

CN Propanamide, N-[3-[1-[6-(3-fluorophenyl)-6-[2-fluoro-5-(trifluoromethyl)phenoxy]hexyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-52-3 CAPLUS

CN Propanamide, N-[3-[1-[6-[2-fluoro-5-(trifluoromethyl)phenoxy]-6phenylhexyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-53-4 CAPLUS

CN Propanamide, N-[3-[1-[7-(2-fluorophenyl)-7-[2-fluoro-5-(trifluoromethyl)phenoxy]heptyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-55-6 CAPLUS
CN Propanamide, N-[3-[1-[5-(4-bromophenoxy)-5-(4-fluorophenyl)pentyl]-4piperiddinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-56-7 CAPLUS

CN Propanamide, N-[3-[1-[5-(4-chlorophenoxy)-5-(4-chlorophenyl)pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-57-8 CAPLUS
- CN Propanamide, N-[3-[1-[5-(4-chlorophenoxy)-5-phenylpenty1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-59-0 CAPLUS
- CN Propanamide, N-[3-[1-[5-(4-chloropheny1)-5-(4-fluorophenoxy)penty1]-4-piperidiny1]pheny1]-2-methy1- (CA INDEX NAME)

- RN 487057-60-3 CAPLUS
- CN Propanamide, N-[3-[1-[5-(4-bromophenoxy)-5-phenylpentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-62-5 CAPLUS
- CN Propanamide, N-[3-[1-[5-(4-fluorophenoxy)-5-(4-fluorophenyl)pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-65-8 CAPLUS

CN Propanamide, N-[3-[1-[5-[2-fluoro-5-(trifluoromethyl)phenoxy]-5-[4-(trifluoromethyl)phenyl]pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-66-9 CAPLUS
- CN Propanamide, N-[3-[1-[5-(3-chlorophenyl)-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

- RN 487057-67-0 CAPLUS
- CN Propanamide, N-[3-[1-[5-(2-fluorophenyl)-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methyl-(CA INDEX NAME)

- RN 487057-68-1 CAPLUS
- CN Propanamide, N-[3-[1-[5-(3-fluorophenyl)-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-70-5 CAPLUS

CN Propanamide, N-[3-[1-[5-(4-chlorophenyl)-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-71-6 CAPLUS

CN Propanamide, N-[3-[1-[5-(4-fluorophenyl)-5-[2-fluoro-5-(trifluoromethyl)phenoxy[pentyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-72-7 CAPLUS

CN Propanamide, N-[3-[1-[5-(4-bromophenoxy)-5-(4-chlorophenyl)pentyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-73-8 CAPLUS

CN Propanamide, N-[3-[1-[5-(4-chlorophenoxy)-5-(4-fluorophenyl)pentyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 487057-76-1 CAPLUS

CN Propanamide, N-[3-[1-[5-(4-fluorophenoxy)-5-phenylpentyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

RN 488098-61-9 CAPLUS

CN Propanamide, N-[3-[1-[6-(4-fluorophenoxy)-6-(2-fluorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

HC1

RN 488098-62-0 CAPLUS

<12/04/2007>

CN Propanamide, N-[3-[1-[6-(4-fluorophenoxy)-6-(4-fluorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

● HC1

RN 488098-63-1 CAPLUS
CN Propanamide, N-[3-[1-[6-(4-chlorophenoxy)-6-(4-fluorophenyl)hexyl]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

- RN 488098-64-2 CAPLUS
- CN Propanamide, N-[3-[1-[6-(4-chlorophenyl)-6-(4-fluorophenoxy)hexyl]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

HC1

RN 488098-65-3 CAPLUS

<12/04/2007>

CN Propanamide, N-[3-[1-[6-(4-chlorophenoxy)-6-phenylhexy1]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

$$\stackrel{\text{Ph}}{\underset{\text{$i$-Pr-C-NH}}{\circ}} \text{C1}$$

● HCl

- RN 488098-67-5 CAPLUS
- CN Propanamide, N-[3-[1-[6-(4-fluorophenoxy)-6-phenylhexyl]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

● HCl

- RN 488098-69-7 CAPLUS
- CN Propanamide, N-[3-[1-[6-(2-fluorophenyl)-6-[2-fluoro-5-(trifluoromethyl)phenoxy]hexyl]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

HC1

- RN 488098-70-0 CAPLUS
- CN Propanamide, N-[3-[1-[5-(4-bromophenoxy)-5-(4-fluorophenyl)pentyl]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

10/513699

HCl

- RN 488098-71-1 CAPLUS
  CN Propanamide, N-[3-[1-[5-(4-chlorophenoxy)-5-(4-chlorophenyl)pentyl]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)
- C1 O-CH-(CH<sub>2</sub>)<sub>4</sub>-N NH-C-Pr-

HCl

RN 488098-72-2 CAPLUS

- CN Propanamide, N-[3-[1-[5-(4-chlorophenoxy)-5-phenylpentyl]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)
- Ph C1

HC1

- RN 488098-73-3 CAPLUS
- CN Propanamide, N-[3-[1-[5-(4-chloropheny1)-5-(4-fluorophenoxy)penty1]-4-piperidiny1]pheny1]-2-methy1-, hydrochloride (1:1) (CA INDEX NAME)

#### HC1

- 488098-74-4 CAPLUS RN
- Propanamide, N-[3-[1-[5-(4-bromophenoxy)-5-phenylpentyl]-4-CN piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

# ● HCl

- RN 488098-76-6 CAPLUS
- CN Propanamide, N-[3-[1-[5-(4-fluorophenoxy)-5-(4-fluorophenyl)pentyl]-4piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

#### HC1

- RN
- 488098-77-7 CAPLUS Propanamide, N-[3-[1-[5-(3-chloropheny1)-5-[2-fluoro-5-CN (trifluoromethy1)phenoxy]penty1]-4-piperidiny1]pheny1]-2-methy1-, hydrochloride (1:1) (CA INDEX NAME)

10/513699

● HCl

RN 488098-78-8 CAPLUS

CN Propanamide, N-[3-[1-[5-(2-fluorophenyl)-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methylhydrochloride (1:1) (CA INDEX NAME)

● HC1

RN 488098-79-9 CAPLUS

CN Propanamide, N-[3-[1-[5-(3-fluoropheny])-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

HC1.

RN 488098-81-3 CAPLUS

<12/04/2007>

Erich Leese

CN Propanamide, N-[3-[1-[5-(4-chloropheny])-5-[2-fluoro-5-(trifluoromethyl)phenoxy]pentyl]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

• HCl

- RN 488098-82-4 CAPLUS
- CN Propanamide, N-[3-[1-[6-(4-fluoropheny1)-6-[2-fluoro-5-(trifluoromethyl)]phenoxy]hexyl]-4-piperidinyl]phenyl]-2-methyl-, hydrochloride (1:1) (CA INDEX NAME)

HCl

REFERENCE COUNT:

3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 14 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2002:31619 CAPLUS

DOCUMENT NUMBER: 136 - 96697

TITLE: Human melanin concentrating hormone receptor MCH1, its

DNA, its synthetic ligands and diagnostic and

therapeutic uses thereof

INVENTOR(S): Salon, John A.; Laz, Thomas M.; Nagorny, Raisa; Wilson, Amv E.

PATENT ASSIGNEE(S):

Synaptic Pharmaceutical Corporation, USA PCT Int. Appl., 524 pp. SOURCE:

CODEN: PIXXD2

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY	ACC.	NUM.	COUNT:	
PATENT	INFO	RMATI	: NC	

	TENT :				KIN	D	DATE								-	ATE	
						-											
	2002						2002			WO 2	001-	US21:	350		2	0010	705
WO	2002	0027	44		A3		2002	8080									
	W:	ΑE,	AG,	AL,	AM,	AT,	ΑU,	ΑZ,	BA,	BB,	BG,	BR,	BY,	ΒZ,	CA,	CH,	CN,
		CO,	CR,	CU,	CZ,	DE,	DK,	DM,	DZ,	EC,	EE,	ES,	FI,	GB,	GD,	GE,	GH,
		GM,	HR,	HU,	ID,	IL,	IN,	IS,	JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LR,
		LS,	LT,	LU,	LV,	MA,	MD,	MG,	MK,	MN,	MW,	MX,	MZ,	NO,	NZ,	PL,	PT,
		RO,	RU,	SD,	SE,	SG,	SI,	SK,	SL,	TJ,	TM,	TR,	TT,	TZ,	UA,	UG,	UZ,
		VN,	YU,	ZA,	ZW												
	RW:	GH,	GM,	KE,	LS,	MW,	MZ,	SD,	SL,	SZ,	TZ,	UG,	ZW,	ΑT,	BE,	CH,	CY,
		DE,	DK,	ES,	FΙ,	FR,	GB,	GR,	ΙE,	IT,	LU,	MC,	NL,	PT,	SE,	TR,	BF,
		ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GW,	ML,	MR,	NE,	SN,	TD,	TG		
CA	2384	358			A1		2002	0110		CA 2	001-	2384	358		2	0010	705
EP	1246	847			A2		2002	1009		EP 2	001-	9524	56		2	0010	705
	R:	AT,	BE,	CH,	DE,	DK,	ES,	FR,	GB,	GR,	IT,	LI,	LU,	NL,	SE,	MC,	PT,
		IE,	SI,	LT,	LV,	FI,	RO,	MK,	CY,	AL,	TR						
JP	2004	5024	23		T		2004	0129		JP 2	002-	5079	86		2	0010	705
PRIORITY	Y APP	LN.	INFO	. :						US 2	000-	6106	35		A 2	0000	705
										WO 2	001-	JS21:	350	1	7 2	0010	705

This invention provides an isolated nucleic acid encoding a human MCH1 receptor, a purified human MCH1 receptor, vectors comprising isolated nucleic acid encoding a human MCH1 receptor, cells comprising such vectors, antibodies directed to a human MCH1 receptor, nucleic acid probes useful for detecting nucleic acid encoding human MCH1 receptors, antisense oligonucleotides complementary to unique sequence of nucleic acid encoding human MCH1 receptors, transgenic, nonhuman animals which express DNA encoding a normal or mutant human MCH1 receptor, methods of isolating a human MCH1 receptor, methods of treating an abnormality that is linked to the activity of a human MCH1 receptor, as well as methods of determining

binding

of compds. to mammalian MCH1 receptors. This invention provides a method of modifying the feeding behavior of a subject which comprises administering to the subject an amount of an MCH1 antagonist effective to decrease the body mass of the subject and/or decrease the consumption of food by the subject. This invention further provides a method of treating a subject suffering from depression and/or anxiety which comprises administering to the subject an amount of an MCH1 antagonist effective to treat the subject's depression and/or anxiety.

387826-68-8P 387826-69-9P 387826-73-5P

<sup>387826-65-5</sup>P 387826-66-6P 387826-67-7P

387826-74-6P 387826-79-1P 387826-80-4P 387826-81-5P 387826-82-6P 387826-85-9P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES

(Uses)
(Numan melanin concentrating hormone receptor MCH1, its DNA, its synthetic ligands and diagnostic and therapeutic uses thereof)

RN 387826-65-5 CAPLUS

CN Propanamide, N-[3-[1-[(3S)-3-(3-chlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 387826-66-6 CAPLUS

CN Propanamide, N-[3-[1-[(3S)-3-(4-chlorophenoxy)-3-phenylpropyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

RN 387826-67-7 CAPLUS

CN Propanamide, 2-methyl-N-[3-[1-[(3S)-3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]-4-piperidinyl]phenyl]- (CA INDEX NAME)

Absolute stereochemistry.

10/513699

RN 387826-68-8 CAPLUS
CN Propanamide, N-[3-[1-[(3R)-3-(2,5-difluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl-2-methyl- (CA INDEX NAME)

### Absolute stereochemistry.

- RN 387826-69-9 CAPLUS
- CN Propanamide, N-[3-[1-[(3R)-3-(3,4-dichlorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

### Absolute stereochemistry.

RN 387826-73-5 CAPLUS

<12/04/2007>

Erich Leese

CN Propanamide, N-[3-[1-[(3\$)-3-(4-fluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-74-6 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-(4-bromophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-79-1 CAPLUS
- CN Propanamide, N-[3-[1-[(3R)-3-[2-fluoro-5-(trifluoromethyl)phenoxy]-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

- RN 387826-80-4 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-[2-fluoro-5-(trifluoromethyl)phenoxy]-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

### Absolute stereochemistry.

- RN 387826-81-5 CAPLUS
- CN Propanamide, N-[3-[1-[(3S)-3-(2,5-difluorophenoxy)-3-phenylpropyl]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

# Absolute stereochemistry.

- RN 387826-82-6 CAPLUS
- CN Propanamide, N-[3-[1-[(3R)-3-(3-chlorophenoxy)-3-phenylpropy1]-4-piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

# Absolute stereochemistry.

RN 387826-85-9 CAPLUS

CN Propanamide, N-[3-[1-[(3S)-3-(2-fluorophenoxy)-3-phenylpropyl]-4piperidinyl]phenyl]-2-methyl- (CA INDEX NAME)

Absolute stereochemistry.

REFERENCE COUNT:

THERE ARE 2 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L3 ANSWER 15 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2001:76 CAPLUS

DOCUMENT NUMBER: 134:207795

TITLE: New 1-aryl-3-(4-arylpiperazin-1-y1)propane

derivatives, with dual action at 5-HT1A serotonin receptors and serotonin transporter, as a new class of

antidepressants

AUTHOR(S): Martinez-Esparza, Javier; Oficialdegui, Ana-M.;

Perez-Silanes, Silvia; Heras, Begona; Orus, Lara; Palop, Juan-A.; Lasheras, Berta; Roca, Joan; Mourelle, Marisa; Bosch, Ana; Del Castillo, Juan-C.; Tordera,

Rosa; Del Rio, Joaquin; Monge, Antonio
CORPORATE SOURCE: Departments of Medicinal Chemistry and Pharmacology

Centro de Investigacion en Farmacobiologia Aplicada (CIFA), Universidad de Navarra, Pamplona, 31080, Spain

SOURCE: Journal of Medicinal Chemistry (2001), 44(3), 418-428

CODEN: JMCMAR; ISSN: 0022-2623
PUBLISHER: American Chemical Society

DOCUMENT TYPE: Journal

LANGUAGE: English

OTHER SOURCE(S): CASREACT 134:207795

GI

AB In a search toward new and efficient antidepressants, l-aryl-3-(4-arylpiperazin-l-yl)propane derivs. I (R = H, Ph, MeO, NO2, Z = CO, CHOH, CHOR1, Rl = 4-F3CC6H4, 4-MeOC6H4, 3,4-OCH3OC6H3, Ar1 =

2-MeOC6H4, 4-ClC6H4, 2-pyridyl, etc.), II (R = H, 2,5-Me2, 5-Me, 5-NO2, Z = CO, CNOH, CHOH, CHOR1, R1 = 4-F3CC6H4, 3,4-OCH2OC6H3, 1-C10H7, position = 2, 3), III and IV (Ar1 = 2-MeOC6H4, 4-C1C6H4, 2-HOC6H4, Z = CO, CHOH) were designed, synthesized, and evaluated for 5-HT reuptake inhibition and 5-HT1A receptor antagonism. This dual pharmacol. profile should lead, in principle, to a rapid and pronounced enhancement in serotoninergic neurotransmission and consequently to a more efficacious treatment of depression. The design was based on coupling structural moieties related to inhibition of serotonin reuptake, such as y-phenoxypropylamines, to arylpiperazines, typical 5-HT1A ligands. In binding studies, several compds. showed affinity at the 5-HT transporter and 5-HT1A receptors. Antidepressant-like activity was initially assayed in the forced swimming test with those compds. with Ki < 200 nM in both binding studies. Functional characterization was performed by measuring the intrinsic effect on rectal temperature in mice and also the antagonism to 8-OH-DPAT-induced hypothermia. The most efficacious compds. II (R = H, Z = CHO-1-C10H7, position = 3, Ar1 = 2-MeOC6H4) (V), II[R = 5-Me, Z = (E)-CNOH, position = 2, Ar1 = 2-MeOC6H4] and IV (Z = CO, CHOH, Ar1 = 2-MeOC6H4) (VI) were further explored for their ability to antagonize 8-OH-DPAT-induced inhibition of forskolin-stimulated cAMP formation in a cell line expressing the 5-HT1A receptor. Furthermore, the antidepressant-like properties of V and VI, which exhibited 5-HT1A receptor antagonistic property in the latter study, were also evaluated in the learned helplessness test in rats. Among these three compds., VI (Z = CHOH) (1-benzo[b]thiophene-3-yl)-3-[4-(2-methoxyphenyl)-1-ylpropan-1-ol] showed the higher affinity at both the 5-HT transporter and 5-HT1A receptors (Ki = 20 nM in both cases) and was also active in the other pharmacol. tests. Such a pharmacol. profile could lead to a new class of antidepressants with a dual mechanism of action and a faster onset of action.

328248-11-9P 328248-15-3P 328248-21-1P ΙT 328248-24-4P 328248-26-6P 328248-30-2P

328248-33-5P 328248-36-8P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(preparation, 5-HT1A serotonin receptor antagonist and serotonin transporter activity, and structure-activity relationship of aryl(arylpiperazinyl)propanes)

RN 328248-11-9 CAPLUS CN

Piperazine, 1-(2-methoxyphenyl)-4-[3-phenyl-3-[4-

(trifluoromethyl)phenoxy[propyl]-, hydrochloride (1:2) (CA INDEX NAME)

$$\stackrel{\text{OMe}}{\underset{N}{\longrightarrow}} \text{N---} \text{CH}_2\text{---} \text{CH}_2\text{---} \text{CH}-\text{O}$$

2 HC1

- RN 328248-15-3 CAPLUS
- CN Piperazine, 1-(4-chlorophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

- RN 328248-21-1 CAPLUS
- CN Piperazine, 1-(4-methoxyphenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

$$\stackrel{\text{Ph}}{\underset{\text{N}}{\longrightarrow}} \text{CF}$$

- RN 328248-24-4 CAPLUS
- CN Piperazine, 1-(2-chlorophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

- RN 328248-26-6 CAPLUS
- CN Piperazine, 1-(4-fluorophenyl)-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]-, hydrochloride (1:2) (CA INDEX NAME)

$$\begin{array}{c} P_h \\ N - CH_2 - CH_2 - CH - O \end{array}$$

### ●2 HC1

- RN 328248-30-2 CAPLUS
- CN Piperazine, 1-(4-nitrophenyl)-4-[3-phenyl-3-[4-

(trifluoromethyl)phenoxy|propyl|- (CA INDEX NAME)

RN 328248-33-5 CAPLUS

CN Piperazine, 1-[3-[1,1'-bipheny1]-4-yl-3-[4(trifluoromethyl)phenoxy[propy1]-4-(2-methoxyphenyl)-, hydrochloride (1:2)
(CA INDEX NABL)

# ●2 HC1

- RN 328248-36-8 CAPLUS
- CN Piperazine, 1-(2-methoxyphenyl)-4-[3-(4-methoxyphenyl)-3-[4-(trifluoromethyl)phenoxy]propyl]- (CA INDEX NAME)

REFERENCE COUNT:

54 THERE ARE 54 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 16 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 2000:601312 CAPLUS

DOCUMENT NUMBER: 133:305272

TITLE: Design, synthesis and biological evaluation of new 3-[(4-aryl)piperazin-1-yl]-1-arylpropane derivatives

> as potential antidepressants with a dual mode of action; serotonin reuptake inhibition and 5-HT1A

receptor antagonism AUTHOR(S):

Oficialdequi, A. M.; Martinez, J.; Perez, S.; Heras, B.; Irurzun, M.; Palop, J. A.; Tordera, R.; Lasheras,

B.; Del Rio, J.; Monge, A.

CORPORATE SOURCE: Department of Medicinal Chemistry, Centro de

Investigacion en Farmacobiologia Aplicada (CIFA), Universidad de Navarra, Pamplona, 31080, Spain

SOURCE . Farmaco (2000), 55(5), 345-353 CODEN: FRMCE8: ISSN: 0014-827X

PUBLISHER: Elsevier Science S.A.

DOCUMENT TYPE: Journal

LANGUAGE: English CASREACT 133:305272

OTHER SOURCE(S):

$$\begin{picture}(0,0) \put(0,0){\line(0,0){100}} \put(0,0){\line(0,0){100}$$

- AB It has been suggested that the combination of a selective serotonin reuptake inhibitor (SSRI) and a 5-HT1A receptor antagonist may facilitate the onset of the SSRIs antidepressant action. Accordingly, we describe the synthesis of a series of new 3-[(4-arvl)piperazin-1-v1]-1-arvlpropane derivs, with structural modifications performed in Arl, Ar2 and Z (Z is different functional groups) to obtain the sought dual activity. Compds. were evaluated for in vitro affinity at 5-HT1A receptors and 5-HT transporter. The antidepressant-like activity of derivs. with the higher affinity was assessed initially using the forced swimming test (FST). Compound 1-(2,4-dimethylphenyl)-3-[(2-methoxyphenyl)piperazin-1-yl]-1propanone (I) showed the best antidepressant-like activity which was further confirmed in the learned helplessness test. 302561-62-2P
- RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(design, synthesis and antidepressant activity of

[(aryl)piperazinyl]arylpropane derivs.)

RN 302561-62-2 CAPLUS CN

Piperazine, 1-(2-methoxyphenyl)-4-[3-(4-methylphenyl)-3-[4-(trifluoromethyl)phenoxy[propyl]- (CA INDEX NAME)

42

REFERENCE COUNT:

THERE ARE 42 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 17 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN

ACCESSION NUMBER: 1994:655827 CAPLUS DOCUMENT NUMBER: 121:255827

ORIGINAL REFERENCE NO.: 121:46707a,46710a

Preparation of (hetero)arylpropanolamine derivatives

as cerebral calcium overload blockers

INVENTOR(S): Jakobsen, Palle; Kanstrup, Anders; Lundbeck, Jane

PATENT ASSIGNEE(S): Novo Nordisk A/S, Den.

Eur. Pat. Appl., 18 pp. SOURCE:

CODEN: EPXXDW DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PATENT NO.	KIND	DATE	APPLICATION NO.	DATE
EP 576766	A1	19940105	EP 1992-610053	19920629
R: GB				
PRIORITY APPLN. INFO.:			EP 1992-610053	19920629

OTHER SOURCE(S):

MARPAT 121:255827 XR3(R0)CCR4R5CR6R7NR1R2 [ X = Ph optionally substituted with one or more cyano, halo, haloalkyl, alkoxy, alkyl, alkanoyl, alkenyl, aryloxy, aralkoxy, amino, alkyl mono or disubstituted amino, alkanoylamino, carbamoyl, alkyl mono- or disubstituted carbamoyl, alkyl substituted with amino, alkyl mono or disubstituted amino, NO2, morpholino, imidazolyl; R = 3,4-methylenedioxyphenyl, aryl or heteroaryl all of which can be optionally substituted with one or more cyano, halogeno, alkyl, alkoxy, alkenyl, trifluoromethyl, alkylene, aryloxy, aralkoxy, alkylthio; R1, R2 = alkyl, cycloalkyl, alkenyl, cycloalkylalkyl, all of which can be unsubstituted or substituted with alkyl, alkoxy or cyano; R1R2 = 5-, 6- or 7-membered ring containing ≥1 N atom, or which optionally contains 2 N atoms, one or 2 O atom(s) or one or 2 S atom(s) or a combination thereof, which ring is optionally substituted with alkyl, alkoxy, or aryl; and R3-R7 = H, alkyl, phenyl; R4X = carbocyclic ring containing 5 or 6 atoms; or salts thereof with a pharmaceutically acceptable acid; with provisos], were prepared Thus, 1-(4-cyanophenyl)-3-piperidinylpropan-1-ol was condensed with 4-trifluoromethylbenzotrifluoride using KOCMe3 to give 1-[3-(4-cyanophenyl)-3-(4-trifluoromethylphenoxy)propyl[piperidine. isolated as the oxalate. The latter inhibited stimulated uptake of 45Ca by rat P2 synaptosomal prepns. with IC50 = 2.2 μg/mL, vs. 26 μg/mL for nifedipine. Generic I formulations are given.

158545-85-8P 158546-06-6P TT

RL: SPN (Synthetic preparation); PREP (Preparation) (preparation of, as cerebral calcium overload blocker)

RN

158545-85-8 CAPLUS

CN Piperazine, 1-phenyl-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]-(CA INDEX NAME)

$$\begin{array}{c|c} & Ph \\ & Ph \\ & Ph \\ \end{array}$$

RN 158546-06-6 CAPLUS

CN Piperazine, 1-phenyl-4-[3-phenyl-3-[4-(trifluoromethyl)phenoxy]propyl]-, ethanedioate (1:1) (CA INDEX NAME)

CM 1

CRN 158545-85-8 CMF C26 H27 F3 N2 O

CM 2

CRN 144-62-7 CMF C2 H2 O4

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=> d hs
'HS' IS NOT A VALID FORMAT FOR FILE 'CAPLUS'
The following are valid formats:
ABS ---- GI and AB
ALL ----- BIB, AB, IND, RE
APPS ----- AI, PRAI
BIB ----- AN, plus Bibliographic Data and PI table (default)
CAN ----- List of CA abstract numbers without answer numbers
CBIB ----- AN, plus Compressed Bibliographic Data
CLASS ----- IPC, NCL, ECLA, FTERM
DALL ----- ALL, delimited (end of each field identified)
DMAX ----- MAX, delimited for post-processing
FAM ----- AN, PI and PRAI in table, plus Patent Family data
FBIB ----- AN, BIB, plus Patent FAM
IND ----- Indexing data
IPC ----- International Patent Classifications
MAX ----- ALL, plus Patent FAM, RE
PATS ----- PI, SO
SAM ----- CC, SX, TI, ST, IT SCAN ----- CC, SX, TI, ST, IT (random display, no answer numbers;
             SCAN must be entered on the same line as the DISPLAY,
             e.g., D SCAN or DISPLAY SCAN)
STD ---- BIB, CLASS
IABS ----- ABS, indented with text labels
IALL ----- ALL, indented with text labels
IBIB ----- BIB, indented with text labels
IMAX ----- MAX, indented with text labels
ISTD ----- STD, indented with text labels
OBIB ----- AN, plus Bibliographic Data (original)
OIBIB ----- OBIB, indented with text labels
SBIB ----- BIB, no citations
SIBIB ----- IBIB, no citations
HIT ----- Fields containing hit terms
HITIND ----- IC, ICA, ICI, NCL, CC and index field (ST and IT)
            containing hit terms
HITRN ----- HIT RN and its text modification
HITSTR ----- HIT RN, its text modification, its CA index name, and
             its structure diagram
HITSEO ----- HIT RN, its text modification, its CA index name, its
            structure diagram, plus NTE and SEQ fields
FHITSTR ---- First HIT RN, its text modification, its CA index name, and
             its structure diagram
FHITSEQ ---- First HIT RN, its text modification, its CA index name, its
             structure diagram, plus NTE and SEQ fields
KWIC ----- Hit term plus 20 words on either side
OCC ----- Number of occurrence of hit term and field in which it occurs
To display a particular field or fields, enter the display field
```

<12/04/2007> Erich Leese

codes. For a list of the display field codes, enter HELP DFIELDS at an arrow prompt (=>). Examples of formats include: TI, TI, AU; BIB, ST, TI, IND; TI, SO. You may specify the format fields in any order and the

information will be displayed in the same order as the format specification.

All of the formats (except for SAM, SCAN, HIT, HITIND, HITRN, HITSTR, FHITSTR, HITSEO, FHITSEO, KWIC, and OCC) may be used with DISPLAY ACC to view a specified Accession Number.

ENTER DISPLAY FORMAT (BIB):

- L3 ANSWER 1 OF 17 CAPLUS COPYRIGHT 2009 ACS on STN
- AN 2009:412461 CAPLUS
- DN 151:496
- TI QSAR study of the 5-HT1A receptor affinities of arylpiperazines using a genetic algorithm-artificial neural network model
- AU Habibi-Yangjeh, Aziz
- CS Department of Chemistry, Faculty of Science, University of Mohaghegh Ardabili, Ardabil, Iran
- SO Monatshefte fuer Chemie (2009), 140(5), 523-530 CODEN: MOCMB7; ISSN: 0026-9247
- PB SpringerWienNewYork
- DT Journal
- LA English
- RE.CNT 46 THERE ARE 46 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

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(FILE 'HOME' ENTERED AT 18:23:43 ON 29 JUN 2009)

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STRUCTURE UPLOADED L1

211 S L1 FULL L2

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FILE 'CAPLUS' ENTERED AT 18:24:36 ON 29 JUN 2009 L3 17 S L2 FULL

=> log y COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION 98.13 284.23 FULL ESTIMATED COST DISCOUNT AMOUNTS (FOR QUALIFYING ACCOUNTS) SINCE FILE TOTAL ENTRY SESSION -13.94 -13.94

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